search research DA. JIANG 554387

=> fil req

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.15 0.15

FULL ESTIMATED COST

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STRUCTURE FILE UPDATES: 13 NOV 2000 HIGHEST RN 302776-13-2 DICTIONARY FILE UPDATES: 13 NOV 2000 HIGHEST RN 302776-13-2

TSCA INFORMATION NOW CURRENT THROUGH July 8, 2000

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Structure search limits have been increased. See HELP SLIMIT for details.

=> e phytostenol/cn 5

E1 1 PHYTOSPHINGOSINE, 1-PHOSPHATE/CN
E2 1 PHYTOSTANOL/CN
E3 0 --> PHYTOSTENOL/CN
E4 1 PHYTOSTEROLIN/CN
E5 1 PHYTOSTEROLS, ETHOXYLATED/CN
=> s e2; d ide can;

L1 1 PHYTOSTANOL/CN

L1 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2000 ACS

RN 127121-08-8 REGISTRY

CN Phytostanol (9CI) (CA INDEX NAME)

MF Unspecified

CI COM, MAN

SR CA

LC STN Files: BIOBUSINESS, BIOSIS, CA, CAPLUS, TOXLIT, USPATFULL

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

6 REFERENCES IN FILE CA (1967 TO DATE)

4 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

6 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 133:149763

REFERENCE 2: 132:127748

REFERENCE 3: 131:184219

REFERENCE 4: 130:68131

130:68131 Prepared by M. Hale 308-4258

```
REFERENCE 5: 125:329276
REFERENCE 6: 113:106467
```

=> e PHYTOSTIMULIN/CN Ε6 1 E7 PHYTOSTREPTIN/CN E8 PHYTOSULFOKINE .ALPHA./CN E9 PHYTOSULFOKINE .BETA./CN E10 PHYTOSULFOKINE-.ALPHA. (ORYZA SATIVA GENE PSK PRECURSOR)/CN E11 PHYTOTON/CN E12 PHYTOTOXIN (MYROTHECIUM RORIDUM)/CN E13 1 PHYTOTOXIN PKZH 1 (VERTICILLIUM DAHLIAE PEPTIDE MOIETY REDUC ED)/CN E14 . PHYTOTOXIN PKZH 1 (VERTICILLIUM DAHLIAE PEPTIDE MOJETY REDUC ED) CYCLIC (6.FWDARW.9)-DISULFIDE/CN E15 PHYTOTOXIN PKZH 1 (VERTICILLIUM DAHLIAE PEPTIDE MOIETY 1 REDUC ED), CYCLIC (6.FWDARW.9)-DISULFIDE/CN PHYTOTOXIN PKZH 1 (VERTICILLIUM DAHLIAE PEPTIDE MOIETY)/CN E16 1 E17 PHYTOTROPINS/CN => e sitostenol/cn 5 E1 1 SITOSTANONE/CN SITOSTANYL P-COUMARATE/CN E2 1 0 --> SITOSTENOL/CN E3 E4 SITOSTEROL/CN 1 SITOSTEROL 3-O-(METHYL .BETA.-D-GLUCURONOPYRANOSIDE)/CN E5

=> s ?sitostenol?/cns

L2 0 ?SITOSTENOL?/CNS

=> s ?sitostanol?/cns

L3 22 ?SITOSTANOL?/CNS

=> s ?phytostenol?/cns

L4 0 ?PHYTOSTENOL?/CNS

=> s ?sitosterol?/cns

L5 174 ?SITOSTEROL?/CNS

=> s ?phytosterol?/cns

L6 5 ?PHYTOSTEROL?/CNS

=> s fatty acids

```
7069 FATTY
          7409 ACIDS
          6120 FATTY ACIDS
L7
                 (FATTY (W) ACIDS)
=> s glyceride/cn
             O GLYCERIDE/CN
L8
=> s ?glyceride?/cns
          1039 ?GLYCERIDE?/CNS
L9
=> s (linoleic acid or linoleate)/cn
             1 LINOLEIC ACID/CN
             1 LINOLEATE/CN
L10
             2 (LINOLEIC ACID OR LINOLEATE)/CN
=> e serum cholesterol/cn 5
E1
                   SERUM AND GLUCOCORTICOID-REGULATED PROTEIN KINASE (MOUSE
GEN
                   E SGK)/CN
                   SERUM AND GLUCOCORTICOID-REGULATED PROTEIN KINASE
(ORYCTOLAG
                   US CUNICULUS STRAIN NEW-ZEALAND-WHITE RENAL CORTEX
COLLECTIN
                   G TUBULE GENE SGK)/CN
             0 --> SERUM CHOLESTEROL/CN
E3
                   SERUM GLOBULIN/CN
E4
             1
                   SERUM GONADOTROPHIN/CN
=> s serum cholesterol
           646 SERUM
             1 SERUMS
            78 SERA
           724 SERUM
                 (SERUM OR SERUMS OR SERA)
          1645 CHOLESTEROL
L11
             O SERUM CHOLESTEROL
               (SERUM (W) CHOLESTEROL)
  fil medl, caplus, biosis, embase, wpids, jicst, conf, ntic,
'NTIC' IS NOT A VALID FILE NAME
Enter "HELP FILE NAMES" at an arrow prompt (=>) for a list of files
that are available. / If you have requested multiple tiles you can
spedify a corrected file name or you can enter "IGNORE" to continue
accessing the remaining file names entered,
ENTER A/FILE NAME/OR (IGNORE) Intis
COST WWW U.S. DOLLARS
                                                  SINCE FILE
                                                                   TOTAL
                         Prepared by M. Hale 308-4258 ENTRY
                                                                 SESSION
                                                                          Page 3
```

FILE 'MEDLINE' ENTERED AT 15:17:38 ON 14 NOV 2000

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=> s (11 or 13 or 15 or 16 or ?hytoste#ol? or ?sitoste#ol? or ?sitosta#ol?)

extstyle ext

TRUNCATION COMBINATION NOT VALID '?PHYTOSTE#OL?'
'#' TRUNCATION SYMBOL NOT VALID WITHIN 'PHYTOSTE#OL?'
'#' TRUNCATION SYMBOL NOT VALID WITHIN 'PHYTOSTE#OL?'
'#' TRUNCATION SYMBOL NOT VALID WITHIN 'PHYTOSTE#OL?'
The truncation symbol # may be used only at the end of a search term.
To specify a variable character within a word use '!', e.g., 'wom!n'
to search for both 'woman' and 'women'. Enter "HELP TRUNCATION" at an arrow prompt (=>) for more information.

=> s (11 or 13 or 15 or 16 or phytoste#ol? or sitoste#ol? or sitosta#ol?)

'#' TRUNCATION SYMBOL NOT VALID WITHIN 'PHYTOSTE#OL?'

The truncation symbol # may be used only at the end of a search term. To specify a variable character within a word use '\', e.g., 'wom!n' Prepared by M. Hale 308-4258'

Page 4

to search for both 'woman' and 'women'. Enter "HELP TRUNCATION" at an arrow prompt (=>) for more information.

=> s (l1 or l3 or l5 or l6 or phytoste!ol? or sitoste!ol? or sitosta!ol?)

```
1798 FILE MEDLINE
1.12
         11957 FILE CAPLUS
L13
          3883 FILE BIOSIS
L14
L15
          2534 FILE EMBASE
LEFT TRUNCATION IGNORED FOR '?SITOSTANOL?' FOR FILE 'WPIDS'
```

LEFT TRUNCATION IGNORED FOR '?SITOSTEROL?' FOR FILE 'WPIDS'

LEFT TRUNCATION IGNORED FOR '?PHYTOSTEROL?' FOR FILE 'WPIDS'

1.16 563 FILE WPIDS

1.17 242 FILE JICST-EPLUS

'CN' IS NOT A VALID FIELD CODE 'CNS' IS NOT A VALID FIELD CODE

LEFT TRUNCATION IGNORED FOR '?SITOSTANOL?' FOR FILE 'CONF'

LEFT TRUNCATION IGNORED FOR '?SITOSTEROL?' FOR FILE 'CONF'

LEFT TRUNCATION IGNORED FOR '?PHYTOSTEROL?' FOR FILE 'CONF' L18

1 FILE CONF 'CN' IS NOT A VALID FIELD CODE

'CNS' IS NOT A VALID FIELD CODE

LEFT TRUNCATION IGNORED FOR '?SITOSTANOL?' FOR FILE 'NTIS'

LEFT TRUNCATION IGNORED FOR '?SITOSTEROL?' FOR FILE 'NTIS'

LEFT TRUNCATION IGNORED FOR '?PHYTOSTEROL?' FOR FILE 'NTIS'

L19 11 FILE NTIS

TOTAL FOR ALL FILES

20989 (L1 OR L3 OR L5 OR L6 OR PHYTOSTE!OL? OR SITOSTE!OL? OR L20 SITOSTA!

Left truncation is not valid in the specified search field in the specified file. The term has been searched without left truncation. Examples: '?TERPEN?' would be searched as 'TERPEN?' and '?FLAVONOID' would be searched as 'FLAVONOID.'

If you are searching in a field that uses implied proximity, and you used a truncation symbol after a punctuation mark, the system may interpret the truncation symbol as being at the beginning of a term. Implied proximity is used in search fields indexed as single words, for example, the Basic Index.

```
=\frac{1}{3} s 20 and (17 or fat/ty)
                            acids or d2.241.81.407/ct or d10.516.251/ct)
     --∕-User BrÆak
SEARCH ENDED BY USER
```

=> s 120 and (17 or fatty acids or d2.241.81.407/ct or d10.516.251/ct)

```
L21
           166 FILE MEDLINE
L22
          2099 FILE CAPLUS
L23
            376 FILE BIOSIS
L24
             96 FILE EMBASE
L25
             52 FILE WPIDS
L26
            29 FILE JICST-EPLUS
```

'CT' IS NOT A VALID FIELD CODE Prepared by M. Hale 308-4258

Phytoste

```
L27
             0 FILE CONF
             3 FILE NTIS
L28
TOTAL FOR ALL FILES
L29
          2821 L20 AND (L7 OR FATTY ACIDS OR D2.241.81.407/CT OR
D10.516.251/CT
=> s 129 and (glyceride? or 19 or d10.516.351/ct or monoacylglycerols)
L30
             9 FILE MEDLINE
L31
           429 FILE CAPLUS
L32
            31 FILE BIOSIS
L33
             4 FILE EMBASE
LEFT TRUNCATION IGNORED FOR '?GLYCERIDE?' FOR FILE 'WPIDS'
L34
             6 FILE WPIDS
L35
              5 FILE JICST-EPLUS
'CNS' IS NOT A VALID FIELD CODE
'CT' IS NOT A VALID FIELD CODE
LEFT TRUNCATION IGNORED FOR '?GLYCERIDE?' FOR FILE 'CONF'
             0 FILE CONF
'CNS' IS NOT A VALID FIELD CODE
LEFT TRUNCATION IGNORED FOR '?GLYCERIDE?' FOR FILE 'NTIS'
             O FILE NTIS
TOTAL FOR ALL FILES
           484 L29 AND (GLYCERIDE? OR L9 OR D10.516.351/CT OR
MONOACYLGLYCEROLS
Left truncation is not valid in the specified search field in the
specified file. The term has been searched without left truncation.
Examples: 'ZTERPEN?' would be searched as TERPEN?' and '?FLAVONOID'
would be searched as 'FLAVONOID.'
If you are searching in a field that uses implied proximity, and you
used a truncation symbol after a planctuation mark, the system may
interpret the truhcation symbol as being at the beginning of a term.
Implied proximity is used in search fields indexed as single words,
for example, the Basic Index.
=> s 138 and (carboxylic(a)acid? or d2.241/ct)
             O FILE MEDLINE
L39
L40
            15 FILE CAPLUS
L41
             1 FILE BIOSIS
L42
             O FILE EMBASE
L43
             1 FILE WPIDS
L44
              4 FILE JICST-EPLUS
'CT' IS NOT A VALID FIELD CODE
L45
             O FILE CONF
             O FILE NTIS
L46
TOTAL FOR ALL FILES
            21 L38 AND (CARBOXYLIC(A) ACID? OR D2.241/CT)
=> s 147 and (linoleic acid or 110 or linoleate or (d2.241.81.436.390 or d10.516.251.355.432 or d18.576.251.355.376.515)%cf258 Page 6
```

```
4 FILE CAPLUS
1.49
L50
             0 FILE BIOSIS
L51
             O FILE EMBASE
L52
             O FILE WPIDS
L53
             2 FILE JICST-EPLUS
'CN' IS NOT A VALID FIELD CODE
'CT' IS NOT A VALID FIELD CODE
L54
             0 FILE CONF
'CN' IS NOT A VALID FIELD CODE
L55
             O FILE NTIS
TOTAL FOR ALL FILES
             6 L47 AND (LINOLEIC ACID OR L10 OR LINOLEATE OR
(D2.241.81.436.390
                OR D10.516.251.355.432 OR D10.516.251.355.310.515)/CT)
=> dup rem 156
DUPLICATE IS NOT AVAILABLE IN 'CONF'.
ANSWERS FROM THESE FILES WILL BE CONSIDERED UNIQUE
PROCESSING COMPLETED FOR L56
L57
              6 DUP REM L56 (O DUPLICATES REMOVED)
=> d 1-6 cbib abs hit
    ANSWER 1 OF 6 CAPLUS COPYRIGHT 2000 ACS
              Document No. 132:77784 Avocado oil. Evolution of lipid
1999:750672
     components during ripening of the fruits of some cultivars grown in
     Southern Italy. Poiana, M.; Giuffre, A. M.; Mincione, B.; Giuffre, F.
     (Istituto di Microbiologia e Tecnologia Agraria e Forestale, Universita
     degli studi di Reggio Calabria, Italy). Riv. Ital. Sostanze Grasse,
     76(6), 257-275 (Italian) 1999. CODEN: RISGAD. ISSN: 0035-6808.
     Publisher: Stazione Sperimentale per le Industrie degli Oli e dei Grassi.
     The ripening of avocado fruits of the Bacon, Hass, and Reed cultivars
AB
     grown in 3 areas of Southern Italy was studied. The oil produced from
     fruits harvested in several stages of ripening and its biol. stability
     were studied. The fruit pulp chem. compn. was examd. and the extd. oil
     was analyzed. The compn. of triglycerides, fatty acids
     , sterols, and tocopherols was detd. The Bacon and Hass cultivars showed
     good oil yields and good oil stability.
AB
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     was analyzed. The compn. of triglycerides, fatty acids
     , sterols, and tocopherols was detd. The Bacon and Hass cultivars showed
     good oil yields and good oil stability.
ST
     avocado ripening oil glyceride fatty acid sterol tocopherol
ΙT
     Fats and Glyceridic oils, analysis
     RL: AMX (Analytical matrix); ANST (Analytical study)
        (avocado; glyceride, fatty acid, sterol and tocopherol compn.
        of avocado oil in relation to fruit ripening stage in 3 cultivars
grown
                         Prepared by M. Hale 308-4258
```

Page 7

O FILE MEDLINE

L48

```
in Southern Italy)
     Growth and development, plant
ΙT
        (fruit ripening; glyceride, fatty acid, sterol and tocopherol
        compn. of avocado oil in relation to fruit ripening stage in 3
        cultivars grown in Southern Italy)
TΤ
    Avocado
        (glyceride, fatty acid, sterol and tocopherol compn. of
        avocado oil in relation to fruit ripening stage in 3 cultivars grown
in
        Southern Italy)
     Carboxylic acids, biological studies
TT
     Carotenes, biological studies
     Chlorophylls, biological studies
     Fatty acids, biological studies
     Glycerides, biological studies
     Lipids, biological studies
     Mineral elements, biological studies
     Sterols
     Tocopherols
     RL: FFD (Food or feed use); BIOL (Biological study); USES (Uses)
        (glyceride, fatty acid, sterol and tocopherol compn. of
        avocado oil in relation to fruit ripening stage in 3 cultivars grown
in
        Southern Italy)
ΙT
     Oxidation
        (lipid; glyceride, fatty acid, sterol and tocopherol compn.
        of avocado oil in relation to fruit ripening stage in 3 cultivars
grown
        in Southern Italy)
IT
     Carbohydrates, biological studies
     RL: FFD (Food or feed use); BIOL (Biological study); USES (Uses)
        (reducing sugars; glyceride, fatty acid, sterol and
        tocopherol compn. of avocado oil in relation to fruit ripening stage
in
        3 cultivars grown in Southern Italy)
TΥ
     57-10-3, Hexadecanoic acid, biological studies
                                                       57-11-4, Octadecanoic
     acid, biological studies
                                57-88-5, Cholesterol, biological studies
     59-02-9, .alpha. Tocopherol 60-33-3, Linoleic
     acid, biological studies 83-45-4, Sitostanol
     83-46-5, .beta.-Sitosterol 83-48-7, Stigmasterol
     112-80-1, Oleic acid, biological studies
                                                112-85-6, Behenic acid
     122-32-7, Triolein
                          148-03-8, .beta. Tocopherol
                                                         373-49-9
     Palmitoleic acid
                        463-40-1, Linolenic acid
                                                    474-60-2, Campestanol
     474-62-4, Campesterol
                             474-63-5, 24-Methylenecholesterol
                                                                  506-12-7.
                          506-30-9, Arachic acid
     Heptadecanoic acid
                                                    516-78-9,
     .DELTA.7-Campesterol 537-40-6
                                     544-63-8, Tetradecanoic acid,
     biological studies 555-44-2, Tripalmitin
                                                557-59-5, Lignoceric
            2364-23-0, Clerosterol
                                      6869-99-4, .DELTA.7-Stigmastenol
     7616-22-0, .gamma. Tocopherol 14465-68-0
                                                 18472-36-1,
     .DELTA.5-Avenasterol 20246-55-3
                                       23290-26-8,
     .DELTA.7-Avenasterol 26836-30-6 26836-35-1
     26836-36-2 26836-37-3 27071-84-7
                                         28040-00-8
     28409-91-8 28409-94-1 28630-67-3
     28880-78-6
                  28933-89-3, Eicosenoic acid
                                                 28949-66-8
     29589-86-4 29661-35-6
                             38703-17-2
                                           38703-23-0
     82181-49-5 87973-00-0 125547-89-9 125547-91-3 Prepared by M. Hale 308-4258
```

RL: FFD (Food or feed use); BIOL (Biological study); USES (Uses) (glyceride, fatty acid, sterol and tocopherol compn. of avocado oil in relation to fruit ripening stage in 3 cultivars grown

in

Southern Italy)

L57 ANSWER 2 OF 6 JICST-EPlus COPYRIGHT 2000 JST
900402115 Chemical composition of lipids, especially triacylglycerol, in grape
seeds.. OHNISHI M; ITO S; FUJINO Y; HIROSE S; KAWAGUCHI M. Obihiro Univ.
Agriculture and Veterinary Medicine, Obihiro, JPN; Tokachi-Ikeda Research
Inst. Viticulture and Enology, Hokkaido, JPN. Agric Biol Chem. (1990)
vol. 54, no. 4, pp. 1035-1042. Journal Code: G0021A (Fig. 1, Tbl. 9, Ref.
24) CODEN: ABCHA6; CODEN: 0002-1369; Pub. Country: Japan. Language:
English.

AB Total lipids were extracted from five varieties of grape seeds and systematically analyzed for their chemical compositions. The yields of the

total lipids were 10-16%, and triacylglycerol(TG) usually amounted to c. 90% of the whole. From a reversed-phase high-performance liquid chromatographic analysis, the major molecular species of TG were shown to be trilinolein (40%), oleoyldilinolein (21%) and palmitoyldilinolein (18%). The component fatty acids were asymmetrically distributed at C-1 and C-3 of the TG molecule. Palmitic acid was exclusively located at the C-1 position, although unsaturated fatty acids, especially linoleic acid

, were predominant at the C-1 position, as at the C-2 and C-3 positions. Compared with TG, higher proportions of palmitic and linolenic acids were generally observed in thirteen other lipid classes isolated from grape seeds, although the fatty acid compositions of the diacylglycerol and

free

fatty acids were roughly identical with that of TG. As
component sterols, sitosterol, campesterol and stigmasterol,
especially the former, were predominant. Their relative proportions were
somewhat different from each other between the neutral and polar sterol
lipids. (author abst.)

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Prepared by M. Hale 308-4258

Page 9

CTgrape; seed; triglyceride; fatty acid composition; breed specificity; vegetable fats and oils; wine making; vitamin F; polyene; aliphatic carboxylic acid; unsaturated carboxylic

acid; olefin compound; diene

BT edible fruit; garden crop; crop(agriculture); agricultural food; food; Vitaceae; Choripetalae; Dicotyledoneae; Angiospermae; Phanerogamae; plant(organism); plant organ; glyceride; carboxylate(ester); ester; lipid; aliphatic alcohol; alcohol; hydroxy compound; chemical composition; composition(constitution); biological comparison; comparison;

fats and oils; oils; fermented food production; food processing; working and processing; fatty acid; carboxylic acid;

fat-soluble vitamin; vitamin

L57 ANSWER 3 OF 6 JICST-EPlus COPYRIGHT 2000 JST 900578062 Compositions of lipid classes, fatty acids and sterols in domestic rye grains.. MANO YASUO; ONISHI MASAO; SATO HARUHIKO; NAKANISHI HAJIME; MAEMOTO MASAMICHI; ITO SEISUKE. Obihirootanitankidaigaku; Obihiro Univ. of Agriculture and Veterinary Medicine. Nippon Shokuhin Kogyo Gakkaishi (Journal of Japanese Society of

Food Science and Technology). (1990) vol. 37, no. 5, pp. 338-345. Journal Code: F0895A (Tbl. 7, Ref. 21) CODEN: NSKGAX; CODEN: 0029-0394; Pub. Country: Japan. Language: Japanese.

Total lipids were extracted from four varieties of domestic rye grains, AB and their lipid class compositions, the component fatty acids and sterols were investigated. The yields of the total lipids were 1.6-1.9%. The neutral lipid fraction usually amounted to about

70% of the whole, in which triacylglycerol was predominant. The ratio of the glycolipid and phospholipid fractions was approximately 1:1.4; the principal lipid classes were diglycosyldiacylglycerol, monoglycosyldiacylglycerol and cerebroside in the former group, and phosphatidylcholine, phosphatidylethanolamine, phosphatidylinositol and lysophosphatidylcholine in the latter one. The main fatty acids in nine lipid classes were generally linoleic, palmitic, oleic and linolenic acids in the decreasing order, except for acylsterol, in which the most abundant one was palmitic acid. The relative proportions

of linoleic acid were 55-60% in triacylglycerol, 65-79% in two glyceroglycolipids and 41-68% in phospholipid classes. Seven

types of 4-desmethylsterols were detected, among which sitosterol and campesterol, particularly the former one, were predominant in neutral and polar sterol lipids. No significant differences were recognized in the

chemical compositions of the rye grain lipids among the varieties harvested in Hokkaido, Japan. (author abst.)

ΤI Compositions of lipid classes, fatty acids and sterols in domestic rye grains.

Total lipids were extracted from four varieties of domestic rye grains, AB and their lipid class compositions, the component fatty acids and sterols were investigated. The yields of the total lipids were 1.6-1.9%. The neutral lipid fraction usually amounted to about

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the

- chemical compositions of the rye grain lipids among the varieties harvested in Hokkaido, Japan. (author abst.)
- CT Secale cereale; Hokkaido; cultivar; lipid; fatty acid composition; plant sterol; triglyceride; aliphatic carboxylic acid; rye flour
- BT cereal; agricultural food; food; common crop; crop(agriculture); Secale; Gramineae; Monocotyledoneae; Angiospermae; Phanerogamae; plant(organism); Japan; East Asia; Asia; breed; chemical composition; composition(constitution); sterol; steroid; derived lipid; glyceride; carboxylate(ester); ester; aliphatic alcohol; alcohol; hydroxy compound; carboxylic acid; cereal flour; processed cereal product
- L57 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2000 ACS
- 1990:213906 Document No. 112:213906 Lipids of oleaster fruits. Goncharova, N. P.; Glushenkova, A. I. (Inst. Khim. Rastit. Veshchestv, Tashkent, USSR). Khim. Prir. Soedin. (1), 17-21 (Russian) 1990. CODEN: KPSUAR. ISSN: 0023-1150.
- AB Lipids were examd. in the seeds and pericarp of 3 morphol. forms of oleaster (Elaeagnus angustifolia). Linoleic acid was the major fatty acid in the seed oil (.ltoreq.59.1% of the triglyceride fraction). Principal fatty acids in the pericarp were palmitic, oleic, and linoleic. .beta.-Sitosterol was the major sterol. Nonacosane accounted for >55% of the alkanes in both seeds and pericarp.
- AB Lipids were examd. in the seeds and pericarp of 3 morphol. forms of oleaster (Elaeagnus angustifolia). Linoleic acid was the major fatty acid in the seed oil (.ltoreq.59.1% of the triglyceride fraction). Principal fatty acids in the pericarp were palmitic, oleic, and linoleic. .beta.-Sitosterol was the major sterol. Nonacosane accounted for >55% of the alkanes in both seeds and pericarp.
- IT Alkanes, biological studies
 Fatty acids, biological studies
 Glycerides, biological studies
 Glycolipids
 Lipids, biological studies
 RL: BIOL (Biological study)
 (of oleaster fruits and seeds)
- Triterpenes and Triterpenoids Prepared by M. Hale 308-4258

(Occurrence) (carboxylic acids, of oleaster fruits) ΙT Glycerides, biological studies RL: BOC (Biological occurrence); BIOL (Biological study); OCCU (Occurrence) (epoxy, of oleaster seeds) TΤ Fatty acids, esters RL: BIOL (Biological study) (esters, of oleaster fruits and seeds) ΙT Carboxylic acids, biological studies RL: BOC (Biological occurrence); BIOL (Biological study); OCCU (Occurrence) (triterpenoid, of oleaster fruits) IT 57-10-3, Hexadecanoic acid, biological studies 57-11-4, C18:0, biological studies 60-33-3, 9,12-Octadecadienoic acid <math>(Z,Z)-, biological studies 112-80-1, 9-Octadecenoic acid (Z)-, biological 143-07-7, Dodecanoic acid, biological studies studies 506-12-7, Heptadecanoic acid Decanoic acid 544-63-8, Tetradecanoic acid, biological studies 1002-84-2, Pentadecanoic acid 28039-99-8 RL: BIOL (Biological study) (of oleaster fruit and seeds) ΙT 112-95-8, Eicosane 593-45-3, Octadecane 593-49-7, 629-92-5, Nonadecane 629-94-7, Heneicosane 629-97-0, Heptacosane 629-99-2, Pentacosane 630-01-3, Hexacosane 630-02-4, Docosane 630-03-5, Nonacosane 630-04-6, Hentriacontane Octacosane 638-67-5, 638-68-6, Triacontane Tricosane 646-31-1, Tetracosane 2363-71-5, 2433-96-7, Tricosanoic acid Heneicosanoic acid 28929-01-3 31152-46-2 RL: BOC (Biological occurrence); BIOL (Biological study); OCCU (Occurrence) (of oleaster fruits) L57 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2000 ACS Document No. 101:188239 Effects of sour rot on the composition of the lipid fraction of different parts of grape berries (Vitis vinifera cv. Fortana). Zironi, R.; Frega, N.; Conte, L. S.; Lercker, G. (Cent. Ric. Vitic. Enol., Univ. Bologna, Bologna, 40126, Italy). Vitis, 23(2), 93-105 (French) 1984. CODEN: VITIAY. ISSN: 0042-7500. AB As compared with healthy grape berries of V. vinifera cv. Fortana, the berries infected with sour rot showed a higher lipid content of the pulp including the skin and a lower lipid content of the seeds. In the seeds from healthy and diseased berries, no difference in fatty acid compn. was obsd., whereas the pulp and skin from infected berries had a higher content of fatty acids with <18 C atoms than did the pulp of healthy berries. An increased oleic and a decreased linoleic acid content was found in the pulp of infected as compared to that of healthy berries. The former showed also a higher ratio between unsatd. and satd. fatty acids. In the pulp and skin oleanolic aldehyde, oleanolic acid, and erythrodiol were detected. The level of the latter was higher in pulp from infected than from healthy berries. AB As compared with healthy grape berries of V. vinifera cv. Fortana, the berries infected with sour rot showed a higher lipid content of the pulp

including the skin and a lower lipid content of the seeds. In the seeds Prepared by M. Hale 308-4258 Page 12

RL: BOC (Biological occurrence); BIOL (Biological study); OCCU

from healthy and diseased berries, no difference in fatty acid compn. was obsd., whereas the pulp and skin from infected berries had a higher content of fatty acids with <18 C atoms than did the pulp of healthy berries. An increased oleic and a decreased linoleic acid content was found in the pulp of infected as compared to that of healthy berries. The former showed also a higher ratio between unsatd. and satd. fatty acids. In the pulp and skin oleanolic aldehyde, oleanolic acid, and erythrodiol were detected. The level of the latter was higher in pulp from infected than from healthy berries.

IT Fatty acids, biological studies Glycerides, biological studies Lipids, biological studies RL: BIOL (Biological study)

(of grapevine parts, sour rot disease effect on)

IT Terpenes and Terpenoids, biological studies

RL: BIOL (Biological study)

(carboxylic acids, of grapevine parts in sour disease)

IT Alcohols, biological studies

Carboxylic acids, biological studies

RL: BIOL (Biological study)

(terpenoid, of grapevine parts in sour disease)

IT **83-46-5** 83-48-7 474-62-4 545-46-0 545-48-2 6869-99-4 17605-67-3 18472-36-1

RL: BIOL (Biological study)

(of grape vine parts in sour rot disease)

IT 57-10-3, biological studies 57-11-4, biological studies 57-88-5,
 biological studies 60-33-3, biological studies 111-02-4
 112-80-1, biological studies 112-85-6 143-07-7, biological studies
 373-49-9 463-40-1 506-12-7 506-30-9 506-38-7 506-46-7

506-48-9

506-50-3 508-02-1 511-61-5 544-63-8, biological studies 544-64-9 559-70-6 557-59-5 1449-09-8 2363-71-5 2433-96-7 3625-52-3 4657-58-3 4250-38-8 7138-40-1 17020-22-3 38232-01-8 60485-38-3

RL: BIOL (Biological study)

(of grapevine parts in sour rot disease)

- L57 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2000 ACS
- 1981:617663 Document No. 95:217663 The constituents of the flower of Ligustrum obtusifolium Sieb. et Zucc. Kikuchi, Masao (Tohoku Coll. Pharm., Sendai, 983, Japan). Nippon Nogei Kagaku Kaishi, 55(9), 821-3 (Japanese) 1981. CODEN: NNKKAA.
- AB Steam distn. of 32 g cold ether ext. from 2 kg flowers of L. obstifolium yielded 0.9 g essential oil consisting of 35 mg acids (I), 50 mg phenols (II), 160 mg of a neutral fraction III (III) and 655 mg of a neutralfraction IV (IV) eluted through silica gel chromatog. with hexane and ether, resp. I was detd. on methylation and gas chromatog. (Silicone OV-17, 60-250.degree.) to contain mainly phenylacetic acid (70%) with n-C6-C12 alkanoic and benzoic acids. From II, III, and IV were isolated phenol, o-, m-, and p-cresol, guaiacol, eugenol (65%); C15-C26 n-alkanes; and cis-3-hexen-1-ol, linalool, benzyl alc., phenylethyl alc., and benzaldehyde, resp. From 29 g solid residue of steam distn. were

on silica gel chromatog. (solvent) followed by gas chromatog. 1.45 g Prepared by M. Hale 308-4258 Page 13

```
n-C21-C33 n-alkanes (C6H14); 1.45 g higher fatty acid ester and 1.45 g
    glycerides, 4.35 g palmitic and oleanolic glycerides
     (C6H14-Et2O 9:1); 0.73 g n-C18-C28 alcs. (4:1); 0.37 g lauric, palmitic,
    linoleic acid (4:1); 0.24 g .alpha.- and .beta.-amyrin
     (PhH-EtOAc 9:1); 0.33 g sitosterol (4:1). From hot ag. residue
     of steam distn., were isolated on cooling 0.2 g quercetin and 0.1 g
     kaempferol and on ether extn. and silica gel chromatog. p-coumaric acid
     0.1, caffeic acid 0.1, ferulic acid 0.2, and p-hydroxy-.beta.-phenylethyl
     alc. 0.2 g.
AB
     Steam distn. of 32 g cold ether ext. from 2 kg flowers of L. obstifolium
     yielded 0.9 g essential oil consisting of 35 mg acids (I), 50 mg phenols
     (II), 160 mg of a neutral fraction III (III) and 655 mg of a
     neutralfraction IV (IV) eluted through silica gel chromatog. with hexane
     and ether, resp. I was detd. on methylation and gas chromatog. (Silicone
    OV-17, 60-250.degree.) to contain mainly phenylacetic acid (70%) with
     n-C6-C12 alkanoic and benzoic acids. From II, III, and IV were isolated
     phenol, o-, m-, and p-cresol, guaiacol, eugenol (65%); C15-C26 n-alkanes;
     and cis-3-hexen-1-ol, linalool, benzyl alc., phenylethyl alc., and
    benzaldehyde, resp. From 29 g solid residue of steam distn. were
isolated
     on silica gel chromatog. (solvent) followed by gas chromatog. 1.45 g
     n-C21-C33 n-alkanes (C6H14); 1.45 g higher fatty acid ester and 1.45 g
     glycerides, 4.35 g palmitic and oleanolic glycerides
     (C6H14-Et2O 9:1); 0.73 g n-C18-C28 alcs. (4:1); 0.37 g lauric, palmitic,
     linoleic acid (4:1); 0.24 g .alpha.- and .beta.-amyrin
     (PhH-EtOAc 9:1); 0.33 g sitosterol (4:1). From hot ag. residue
     of steam distn., were isolated on cooling 0.2 g quercetin and 0.1 g
     kaempferol and on ether extn. and silica gel chromatog. p-coumaric acid
     0.1, caffeic acid 0.1, ferulic acid 0.2, and p-hydroxy-.beta.-phenylethyl
     alc. 0.2 q.
IT
     Carboxylic acids, biological studies
     RL: BIOL (Biological study)
        (alkanoic, of Privet flower)
ΙT
     Alcohols, biological studies
     Alkanes, biological studies
     Fatty acids, esters
     Fatty acids, biological studies
     Glycerides, biological studies
     Phenols, biological studies
     RL: BOC (Biological occurrence); BIOL (Biological study); OCCU
     (Occurrence)
        (of Privet flower)
     60-12-8
               65-85-0, biological studies
                                             78-70-6 83-46-5
                                                      100-51-6, biological
               95-48-7, biological studies
                                             97-53-0
               100-52-7, biological studies
                                             103-82-2, biological studies
                                    108-39-4, biological studies
     106-44-5, biological studies
                                                                   108-95-2,
                          117-39-5
     biological studies
                                     331-39-5
                                                501-94-0
                                                           520-18-3
                                                                      559-70-6
     638-95-9
                928-96-1
                           1135-24-6
                                       7400-08-0
     RL: BOC (Biological occurrence); BIOL (Biological study); OCCU
     (Occurrence)
        (of Privet flower)
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=> s fabry b?/au,in

^{&#}x27;IN' IS NOT A VALID FIELD CODE Prepared by M. Hale 308-4258

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18 FILE MEDLINE
L58
          219 FILE CAPLUS
L59
      25 FILE BIOSIS
L60
'IN' IS NOT A VALID FIELD CODE
L61 25 FILE EMBASE
L62
L63
           182 FILE WPIDS
           O FILE JICST-EPLUS
'AU' IS NOT A VALID FIELD CODE
'IN' IS NOT A VALID FIELD CODE
            O FILE CONF
'IN' IS NOT A VALID FIELD CODE
             O FILE NTIS
TOTAL FOR ALL FILES
          469 FABRY B?/AU,IN
=> s 166 and 120
· L67
             O FILE MEDLINE
L68
             5 FILE CAPLUS
             O FILE BIOSIS
             O FILE EMBASE
             4 FILE WPIDS
             O FILE JICST-EPLUS
L73
             0 FILE CONF
L74
             O FILE NTIS
TOTAL FOR ALL FILES
             9 L66 AND L20
=> s 175 not 156
L76
             O FILE MEDLINE
             5 FILE CAPLUS
L78
             0 FILE BIOSIS
L79
             0 FILE EMBASE
L80
            4 FILE WPIDS
L81
             O FILE JICST-EPLUS
L82
             0 FILE CONF
L83
             O FILE NTIS
TOTAL FOR ALL FILES
             9 L75 NOT L56
=> dup rem 184
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DUPLICATE IS NOT AVAILABLE IN 'CONF'. ANSWERS FROM THESE FILES WILL BE CONSIDERED UNIQUE PROCESSING COMPLETED FOR L84 7 DUP REM L84 (2 DUPLICATES REMOVED)

=> d cbib abs 1-7

L85 ANSWER 1 OF 7 CAPLUS COPYRIGHT 2000 ACS DUPLICATE 1 2000:259954 Document No. 132:283924 Use of nanoscale sterols and sterol Prepared by M. Hale 308-4258 Page 15 esters for producing cosmetic and/or pharmaceutical preparations. Foerster, Thomas; Fabry, Bernd; Hollenbrock, Martina; Kropf, Christian (Henkel Kommanditgesellschaft auf Aktien, Germany). PCT Int. Appl. WO 2000021490 A1 20000420, 29 pp. DESIGNATED STATES: W: AU, BG, BR, BY, CA, CN, CZ, HU, ID, IN, IS, JP, KR, LT, LV, MX, NO, NZ, PL, RO, RU, SI, SK, TR, UA, US, ZA; RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE. (German). CODEN: PIXXD2. APPLICATION: WO 1999-EP7359 19991005. PRIORITY: US 1998-PV104144 19981014. When sterols and/or sterol esters with particle diams. of 10-300 nm are used for producing cosmetic and/or pharmaceutical prepns., the fineness

of
the particles compared to known sterol and sterol ester prepns. ensures
that they penetrate into the stratum corneum quickly when applied
topically. Thus, nanoparticulate phytosterols were prepd. by
rapid decompression and expansion of a supercrit. soln. of the sterols in

CO2 at 160.degree. into an expansion chamber contg. a 4% aq. soln. of PEG, followed by evapn. to dryness. A water-in-oil sunscreen cream was prepd. contg. Dehymuls PGPH 2.0, Lameform TGI 4.0, beeswax 3.0, Plantaren 818

5.0, dioctyl carbonate 5.0, Cetiol J600 2.0, Cetiol OE 3.0, panthenol + bisabolol 1.2, **phytosterols** 0.5, Neo Heliopan Hydro 3.0, Neo Heliopan BB 1.5, Neo Heliopan E 1000 5.0, Neo Heliopan AV 4.0, Uvinul T 150 2.0, 86% glycerin 5.0, preservative, and H2O to 100 wt.%.

L85 ANSWER 2 OF 7 CAPLUS COPYRIGHT 2000 ACS

- 1999:344854 Document No. 130:347399 Use of mixtures containing phytostenols for producing hypocholesteremic preparations.

 Fabry, Bernd (Henkel Kommanditgesellschaft auf Aktien, Germany).

 PCT Int. Appl. WO 9925362 A1 19990527, 19 pp. DESIGNATED STATES: W: AU, BG, BR, BY, CA, CN, CZ, HU, ID, IS, JP, KR, LT, LV, MX, NO, NZ, PL, RO, RU, SI, SK, TR, UA, US; RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE. (German). CODEN: PIXXD2. APPLICATION: WO 1998-EP7059 19981105. PRIORITY: DE 1997-19750453 19971114.
- AB Mixts. of active agents contg. (a) phytostenols and/or phytostenol esters and (b) conjugated fatty acids or their glycerides are used to produce hypocholesteremic prepns. These mixts. have a synergistic effect in reducing the cholesterol content of serum. When encapsulated in gelatin, the prepns. can be administered orally in high doses without any problems; they may also be incorporated into food products. Thus, the contents of a 1.5-g capsule, contg. 5 wt.% .beta.-sitostanyl laurate, 5 wt.% conjugated linoleic acid, and radiolabeled cholesterol, were administered to fasting rats by gavage. The radioactivity level in the blood 48 h later was 12% of that in rats fed labeled cholesterol alone, and was also markedly lower than that in rats given either the phytostanol or the fatty acid alone.
- L85 ANSWER 3 OF 7 WPIDS COPYRIGHT 2000 DERWENT INFORMATION LTD
- AN 1999-314061 [27] WPIDS

AΒ

AB DE 19750453 A UPAB: 19990714

NOVELTY - The preparation of a hypocholesterinemic agent (A) comprises mixing: (a) phytostenol and/or phytostenol ester; and

- (b) fatty acids with 6--24C and at least two conjugated double bonds, especially their glycerides.
 - USE (A) is used to lower the cholesterol levels in mammal serum.

- 1998:771359 Document No. 130:25230 Use of selected phytosterol esters for preparation of hypocholesterolemic agents. Fabry, Bernd (Henkel K.-G.a.A., Germany). Ger. DE 19750422 C1 19981126, 6 pp. (German). CODEN: GWXXAW. APPLICATION: DE 1997-19750422 19971114.
- AB Use of **phytosterol** esters of unsatd. conjugated fatty acids for prepn. of hypocholesterolemic agents is described. Thus, in a gelatin capsule is added a mixt. of different .beta.-sitosterol esters (5%), radioactively-labeled cholesterol (0.5%) and if necessary vitamin E.
- The blood of animals receiving these capsules were tested for radioactivity at 3, 6, 12, 24, and 48 h; after 48 h radioactivity was down to 15-21%.
- L85 ANSWER 5 OF 7 CAPLUS COPYRIGHT 2000 ACS
- 1998:385506 Document No. 129:36452 Use of mixtures of phytostanols and tocopherols for the production of hypocholesteremic agents. Weitkemper, Norbert; Fabry, Bernd (Henkel Kommanditgesellschaft Auf Aktien, Germany; Weitkemper, Norbert; Fabry, Bernd). PCT Int. Appl. WO 9823275
- A1

 19980604, 15 pp. DESIGNATED STATES: W: AU, BR, CA, CN, CZ, HU, JP, KR, MX, NO, NZ, PL, RU, SI, SK, US; RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE. (German). CODEN: PIXXD2. APPLICATION: WO 1997-EP6447 19971119. PRIORITY: DE 1996-19649286 19961128; DE 1997-19700796 19970113.
- AB Tocopherols, though themselves having little or no hypocholesteremic activity, potentiate the hypocholesteremic action of phytostanol esters. The effect is further potentiated by chitosan, phytosterol sulfates, RNA, and DNA. If these agents are encapsulated in gelatin, they
 - can be administered orally in high doses without problems. Thus, a gelatin capsule contg. .beta.-sitostanol laurate 5, vitamin E 5, and radiolabeled cholesterol 0.5 wt.% was administered to rats by gavage. The level of blood radioactivity 24 h later was 39% of that in control rats, compared to 51% in rats receiving .beta.-sitostanol laurate but not vitamin E.
- L85 ANSWER 6 OF 7 CAPLUS COPYRIGHT 2000 ACS
- 1998:208621 Document No. 128:235001 Skin care compositions containing esterquats and sterols. Ansmann, Achim; Fabry, Bernd (Henkel K.-G.a.A., Germany). Ger. DE 19652302 Cl 19980326, 8 pp. (German) CODEN: GWXXAW. APPLICATION: DE 1996-19652302 19961216.
- AB Skin-conditioning compns. contg. sterols 0.01-3, oils 1-90, and esterquats
- 0.1-10 wt.% as cationic emulsifiers form oil-in-water emulsions which are stable during storage at elevated temps. Thus, an emulsion contg. Me-quaternized ditallow fatty acid triethanolamine ester methosulfate 5.0,
 - ceteareth-20 5.0, cetearyl glucoside + cetearyl alc. 5.0, phytosterols 1.0, coco glycerides 10.0, oleyl oleate 6.0, almond oil 2.0, 86% glycerin 3.0, and water to 100 wt.% had a viscosity of 20.0 Pa s immediately after prepn. and 19.5 Pa s after 2 days storage at 35.degree..
- L85 ANSWER 7 OF 7 WPIDS COPYRIGHT 2000 DERWENT INFORMATION LTD AN 1998-313690 [28] WPIDS Prepared by M. Hale 308-4258

```
DE 19700796 A UPAB: 19980715
AB
     Use of an active agent mixture (I) for the preparation of
    hypocholesterolaemic agents, is new.
          (I) comprises (A) phytosterols and/or phytosterol
     esters and (B) potentiating agents selected from tocopherols, chitosans,
    phytosterol sulphates and/or (deoxy)ribonucleic acids.
          Also claimed is the use of gelatin for encapsulating (A) or (I).
          USE - (I) is preferably administered orally in gelatin capsules, but
    may also be used in rectal or vaginal suppositories or dissolved or
     dispersed in foodstuffs.
          ADVANTAGE - (B) (which themselves have no hypocholesterolaemic
     activity) potentiate and accelerate the action of (A) in reducing serum
     cholesterol levels.
          Encapsulation of (I) or (A) in gelatin allows oral administration
     without prior art problems of taste and/or consistency.
     Dwq.0/0
=> s (anticholesterem? or hypocholesterem? or (d10.162.202 or
d27.505.519.162.202)/ct)
L86
          5020 FILE MEDLINE
L87
          7670 FILE CAPLUS
L88
            83 FILE BIOSIS
L89
            53 FILE EMBASE
L90
           138 FILE WPIDS
           102 FILE JICST-EPLUS
'CT' IS NOT A VALID FIELD CODE
L92
             0 FILE CONF
L93
             1 FILE NTIS
TOTAL FOR ALL FILES
         13067 (ANTICHOLESTEREM? OR HYPOCHOLESTEREM? OR (D10.162.202 OR
D27.505
               .519.162.202)/CT)
=> s (11 or 13 or 15 or 16 or phytoste!ol? or sitoste!ol? or sitosta!ol?) and
T.95
            85 FILE MEDLINE
L96
           131 FILE CAPLUS
             2 FILE BIOSIS
L97
L98
             1 FILE EMBASE
LEFT TRUNCATION IGNORED FOR '?SITOSTANOL?' FOR FILE 'WPIDS'
LEFT TRUNCATION IGNORED FOR '?SITOSTEROL?' FOR FILE 'WPIDS'
LEFT TRUNCATION IGNORED FOR '?PHYTOSTEROL?' FOR FILE 'WPIDS'
L99
             4 FILE WPIDS
L100
             2 FILE JICST-EPLUS
'CN' IS NOT A VALID FIELD CODE
'CNS' IS NOT A VALID FIELD CODE
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'CNS' IS NOT A VALID FIELD CODE
Prepared by M. Hale 308-4258
                                                                        Page 18
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TOTAL FOR ALL FILES

225 (L1 OR L3 OR L5 OR L6 OR PHYTOSTE!OL? OR SITOSTE!OL? OR L103 SITOSTA!

OL?) AND L94

Left truncation is not valid in the specified search field in the specified file. The term has been searched without left truncation. Examples: '?TERPEN?' would be searched as 'TERPEN?' and '?FLAVONOID' would be searched as 'FLAVONOID.'

If you are searching in a field that uses implied proximity, and you used a truncation symbol after a punctuation mark, the system may interpret the truncation symbol as being at the beginning of a term. Implied proximity is used in search fields indexed as single words, for example, the Basic Index.

=> s 1103 and (treat? or therap? or lower? or reduc?) and serum cholesterol

L104	21	FILE	MEDLINE
L105	22	FILE	CAPLUS
L106	1	FILE	BIOSIS
L107	0	FILE	EMBASE
COMMAND	INTERRU	JPTED	
L108	0	FILE	JICST-EPLUS
L109	0	FILE	CONF
L110	0	FILE	NTIS

TOTAL FOR ALL FILES

L11144 L103 AND (TREAT? OR THERAP? OR LOWER? OR REDUC?) AND SERUM CHOLE

If this message appears repeatedly, please notify the Help Desk. Enter "HELP STN" for information on contacting the nearest STN Help Desk by telephone or via SEND in the STNMAIL file.

=> dup rem 1111

DUPLICATE IS NOT AVAILABLE IN 'CONF'. ANSWERS FROM THESE FILES WILL BE CONSIDERED UNIQUE PROCESSING COMPLETED FOR L111

40 DUP REM L111 (4 DUPLICATES REMOVED)

=> d 1-40 cbib abs

L112 ANSWER 1 OF 40 CAPLUS COPYRIGHT 2000 ACS Document No. 133:17688 Preperation of phytosterol and/or phytostanol derivatives for redn. of serum Prepared by M. Hale 308-4258

cholesterol and triglycerides. Burdick, David Carl; Moine,
Gerard; Raederstorff, Daniel; Weber, Peter (F. Hoffmann-La Roche A.-G.,
Switz.). Eur. Pat. Appl. EP 1004594 Al 20000531, ll pp. DESIGNATED
STATES: R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC,
PT, IE, SI, LT, LV, FI, RO. (English). CODEN: EPXXDW. APPLICATION: EP
1999-122978 19991119. PRIORITY: EP 1998-122412 19981126; EP 1999-119337
19990929.

AB Phytosterol and/or phytostanol esters with polyunsatd. fatty acids having from 18 to 22 carbon atoms and at least three carbon-carbon double bonds are were prepd. as agents effective in reducing both serum cholesterol and triglycerides. Thus, .91 g docosahexaenoic acid was treated with 1.03 g stigmasterol in presence of dimethylaminopyridine in CH2Cl2 to give 1.0 g stigmasterol docosahexaenoate as an oil.

L112 ANSWER 2 OF 40 MEDLINE DUPLICATE 1 2000270456 Document Number: 20270456. Noncholesterol sterols and cholesterol

lowering by long-term simvastatin treatment in coronary
patients: relation to basal serum cholestanol. Miettinen T A; Strandberg

E; Gylling H. (Department of Medicine, University of Helsinki, Helsinki, Finland.. tatu.a.miettinen@helsinki.fi) . ARTERIOSCLEROSIS, THROMBOSIS, AND VASCULAR BIOLOGY, (2000 May) 20 (5) 1340-6. Journal code: B89. ISSN: 1079-5642. Pub. country: United States. Language: English.

AΒ Coronary patients with low baseline ratios of serum cholestanol and plant sterols to cholesterol (indicating low cholesterol absorption) but not those with high ratios (high absorption) experienced reduced recurrences of coronary events during simvastatin treatment in the Scandinavian Simvastatin Survival Study. Thus, in the present study, serum cholesterol, its precursor sterols (reflecting cholesterol synthesis), plant sterols (campesterol and sitosterol), and cholestanol were measured before and during a 5-year period of placebo treatment (n=433) and simvastatin treatment (n=434) in patients from a subgroup of the Scandinavian Simvastatin Survival Study to determine whether changes in cholesterol synthesis and serum levels were related to cholesterol absorption. Serum cholesterol level was unchanged, the ratios of cholesterol precursor sterols to cholesterol were decreased, and the ratios of plant sterols to cholesterol were increased in relation to increasing baseline ratios of cholestanol quartiles. The latter predicted 5-year ratios and simvastatin-induced reductions of the precursor sterols, with the lowering of the ratios (cholesterol synthesis reduction) being almost twice higher in the lowest versus the highest quartile. The ratios of plant sterols, especially campesterol, to cholesterol were markedly increased during simvastatin treatment , mostly in subjects with the highest baseline cholestanol quartiles. Simvastatin reduced serum cholesterol more

(P=0.003) in the lowest versus the highest cholestanol quartile during the

5-year treatment period. The results show for the first time that baseline cholesterol metabolism, measured by serum noncholesterol sterols, predicts the effectiveness of simvastatin in reducing cholesterol synthesis and serum levels of cholesterol. The drug presses

the synthesis of cholesterol markedly more effectively in subjects with Prepared by M. Hale 308-4258 Page 20

high than with low baseline synthesis but reduces respective serum cholesterol levels less markedly than synthesis.

Subjects with high cholesterol absorption and low synthesis may need a combination therapy to lower more effectively their serum cholesterol levels and prevent an increase in the levels of plant sterols.

L112 ANSWER 3 OF 40 MEDLINE

- 2000197829 Document Number: 20197829. Soy sterol esters and betasitostanol ester as inhibitors of cholesterol absorption in human
 small bowel. Normen L; Dutta P; Lia A; Andersson H. (Department of
 Clinical Nutrition, Annedalsklinikerna, Goteborg University, Goteborg,
 Sweden.. nutrition@clinnutr.gu.se). AMERICAN JOURNAL OF CLINICAL
 NUTRITION, (2000 Apr) 71 (4) 908-13. Journal code: 3EY. ISSN: 0002-9165.
 Pub. country: United States. Language: English.
- AB BACKGROUND: Plant sterols are natural dietary components with serum cholesterol-lowering properties. The lowering of serum cholesterol by plant sterols is believed to be the result of an inhibition of cholesterol absorption
- the small bowel, although increased bile acid excretion has also been suggested. The difference in effect of saturated and unsaturated plant sterols on cholesterol absorption needs to be elucidated further.

 OBJECTIVE: The primary aim of this study was to measure small-bowel cholesterol absorption and sterol excretion in addition to hepatic cholesterol synthesis after intake of soy sterol esters and beta-sitostanol ester corresponding to 1.5 g plant sterols/d. DESIGN:

 Seven ileostomy subjects were studied during a control period and 2 intervention periods when either soy sterol esters or beta-sitostanol ester was added to a basal diet. Ileostomy bags were collected every other hour and frozen immediately for analysis of nutrients and sterols. RESULTS: Cholesterol absorption was 56% (43-65%)
- the control period and decreased to 38% (32-46%) in the soy sterol ester period (P = 0.00) and to 39% (30-48%) in the beta-sitostanol ester period (P = 0.00). CONCLUSION: Esterified soy sterols and beta-sitostanol inhibited cholesterol absorption equally, despite the different structures of the plant sterols.

L112 ANSWER 4 OF 40 MEDLINE

2000202391 Document Number: 20202391. Plant stanol esters affect serum cholesterol concentrations of hypercholesterolemic men and women in a dose-dependent manner. Hallikainen M A; Sarkkinen E S; Uusitupa M I. (Department of Clinical Nutrition, University of Kuopio, 70211 Kuopio, Finland.) JOURNAL OF NUTRITION, (2000 Apr) 130 (4) 767-76. Journal code: JEV. ISSN: 0022-3166. Pub. country: United States. Language:

English.

AB The effect of plant stanol ester on **serum cholesterol** is dose-dependent. However, it is not clear what the dose is beyond which no additional benefit can be obtained. Therefore, we determined the dose-response relationship for **serum cholesterol** with different doses of plant stanol ester in hypercholesterolemic subjects.

In a single-blind design each of 22 men or women consumed five different doses of plant stanol [target (actual) intake 0 (0), 0.8 (0.8), 1.6 (1.6), Prepared by M. Hale 308-4258 Page 21

2.4 (2.3), 3.2 (3.0) g/d] added as plant stanol esters to margarine for 4 wk. The order of dose periods was randomly determined. Serum total cholesterol concentration decreased (calculated in reference to control) by 2.8% (P = 0.384), 6.8% (P < 0.001), 10.3% (P < 0.001) and 11.3% (P < 0.001) 0.001) by doses from 0.8 to 3.2 g. The respective decreases for LDL cholesterol were 1.7% (P = 0. 892), 5.6% (P < 0.05), 9.7% (P < 0.001) and 10.4% (P < 0.001). Although the decreases were numerically greater with 2.4 and 3.2 g doses than with the 1.6 g dose, these differences were not significant (P = 0.054-0.516). Serum plant stanols rose slightly, but significantly with the dose (P < 0.001). Apolipoprotein B concentration was decreased significantly already at the dose of 0.8 g (8.7%, P < 0.001). Apolipoprotein E genotype did not affect the lipid responses. We conclude that significant reduction of serum total and LDL cholesterol concentrations is reached with the 1.6-q stanol dose, and increasing the dose from 2.4 to 3.2 g does not provide clinically important additional effect.

L112 ANSWER 5 OF 40 MEDLINE

DUPLICATE 2

2000136131 Document Number: 20136131. Stanol ester margarine alone and with simvastatin lowers serum cholesterol in

families with familial hypercholesterolemia caused by the FH-North Karelia

mutation. Vuorio A F; Gylling H; Turtola H; Kontula K; Ketonen P; Miettinen T A. (Department of Medicine, University of Helsinki, Helsinki, Finland. alpo.vuorio@huch.fi) . ARTERIOSCLEROSIS, THROMBOSIS, AND VASCULAR BIOLOGY, (2000 Feb) 20 (2) 500-6. Journal code: B89. ISSN: 1079-5642. Pub. country: United States. Language: English.

AB In heterozygous familial hypercholesterolemia (FH), serum low density lipoprotein (LDL) cholesterol levels are already elevated at birth. Premature coronary heart disease occurs in approximately 30% of heterozygous untreated adult patients. Accordingly, to retard development of atherosclerosis, preventive measures for lowering cholesterol should be started even in childhood. To this end, 19 FH families consumed dietary stanol ester for 3 months. Stanol ester margarine lowers the serum cholesterol level by inhibiting cholesterol

absorption. Each individual in the study replaced part of his or her daily

dietary fat with 25 g of 80% rapeseed oil margarine containing stanol esters (2.24 g/d stanols, mainly sitostanol). The families who consumed this margarine for 12 weeks included 24 children, aged 3 to 13 years, with the North Karelia variant of FH (FH-NK), 4 FH-NK parents, and 16 healthy family members, and a separate group of 12 FH-NK adults who consumed the margarine for 6 weeks and who were on simvastatin therapy (20 or 40 mg/d). Fat-soluble vitamins were measured by high-pressure liquid chromatography, and cholesterol precursor sterols (indexes of cholesterol synthesis) and cholestanol and plant sterols (indexes of cholesterol absorption efficiency) were assayed by gas-liquid chromatography. No side effects occurred. Serum LDL cholesterol levels were reduced by 18% (P<0.001), 11%, 12% (P<0.001), and 20% (P<0.001) in the 4 groups, respectively. The serum campesterol-tocholesterol ratios fell by 31% (P<0.001), 29%, 23% (P<0.001), and 36% (P<0.001), respectively, suggesting that cholesterol absorption efficiency

was inhibited. Serum lathosterol ratios were elevated by 38% (P<0.001), 11%, 15% (P<0.001), and 19% (P<0.001), respectively, suggesting that cholesterol synthesis was compensatorily upregulated. The FH-NK children Prepared by M. Hale 308-4258

increased their serum lathosterol ratio more than did the FH-NK adults treated with stanol ester margarine and simvastatin (P<0.01). In the FH-NK children, serum retinol concentration and alpha-tocopherol-to-cholesterol ratios were unchanged by stanol ester margarine, but alpha-and beta-carotene concentrations and ratios were decreased. As assayed in a genetically defined population of FH patients, a dietary regimen with stanol ester margarine proved to be a safe and effective hypolipidemic treatment for children and adults. In FH-NK adults on simvastatin therapy, serum LDL cholesterol levels could be reduced even further by including a stanol ester margarine in the regimen.

L112 ANSWER 6 OF 40 MEDLINE
2000002453 Document Number: 20002453. Plant lipids that lower
serum cholesterol [editorial]. Thompson G R. EUROPEAN

HEART JOURNAL, (1999 Nov) 20 (21) 1527-9. Journal code: EM8. ISSN: 0195-668X. Pub. country: ENGLAND: United Kingdom. Language: English.

L112 ANSWER 7 OF 40 CAPLUS COPYRIGHT 2000 ACS

1999:529029 Document No. 131:149329 Composition for reducing serum cholesterol levels. Sorkin, Harlan Lee, Jr.

(USA). PCT Int. Appl. WO 9940922 A1 19990819, 10 pp. DESIGNATED STATES: W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK,

EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL,

PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU,

ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (English). CODEN: PIXXD2. APPLICATION: WO

1999-US3003 19990212. PRIORITY: US 1998-22807 19980212.

AB A compn. for reducing serum cholesterol in humans and animals is provided. The compn. includes phytosterol and policosanol which together produce a synergistic effect in lowering serum cholesterol levels. Preferably the compn. includes about 3.2:1 parts by wt. of phytosterol and policosanol. A tablet contained cholestatin (.gtoreq.88 % phytosterols) 250, rice bran wax (23-33 % policosanol) 250, Ca phosphate 261.7, cellulose 49.4, stearic acid 23.8, Mg stearate 6.8, and silica 9.4 mg.

L112 ANSWER 8 OF 40 CAPLUS COPYRIGHT 2000 ACS

1999:404806 Document No. 131:49483 Sterol esters as food additives.
Milstein, Norman; Biermann, Manfred; Leidl, Peter; Von Kreis, Rainer
(Henkel Corporation, USA). PCT Int. Appl. WO 9930569 Al 19990624, 38 pp.
DESIGNATED STATES: W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH,
CN,

CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (English). CODEN: PIXXD2. APPLICATION:

WO 1998-US26212 19981215. PRIORITY: US 1997-69790 19971216; US 1998-72434

19980504; US 1998-83584 19980521.

AB A food additive useful for lowering serum

cholesterol in humans contains a sterol or stanol ester of a fatty
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acid or a dicarboxylic acid ester of a sterol or stanol made by reacting

sterol, stanol and a carboxylic acid in the presence of an effective amt. of a catalyst selected from the group consisting of calcium oxide, calcium

hydroxide, a calcium salt of a carboxylic acid, magnesium hydroxide and combinations thereof described herein.

L112 ANSWER 9 OF 40 MEDLINE

2000069312 Document Number: 20069312. Effects of low-fat stanol ester enriched margarines on concentrations of serum carotenoids in subjects with elevated **serum cholesterol** concentrations.

Hallikainen M A; Sarkkinen E S; Uusitupa M I. (Department of Clinical Nutrition, University of Kuopio, Kuopio, Finland..

Maarit.Hallikainen@uku.fi) . EUROPEAN JOURNAL OF CLINICAL NUTRITION, (1999

Dec) 53 (12) 966-9. Journal code: EJC. ISSN: 0954-3007. Pub. country: ENGLAND: United Kingdom. Language: English.

AB OBJECTIVE: To investigate the effects of low-fat stanol ester margarines on concentrations of serum carotenoids. DESIGN: A randomized parallel double-blind study design consisting of a 4-week run-in (high-fat diet) and an 8-week experimental (low-fat, low-cholesterol diet) period. During the experimental diet period subjects consumed low-fat wood stanol ester (WSEM), vegetable oil stanol ester (VOSEM) or control (no stanol esters) margarine daily. The daily mean total stanol intake was 2.31 and 2.16 g

the WSEM and VOSEM groups, respectively. SETTING: Outpatient clinical trial with free-living subjects. SUBJECTS: Altogether, 60 hypercholesterolaemic subjects were selected for the study out of 91 originally screened. The study was completed by 55 subjects. MAIN DUTCOMES

MEASURES: Serum alpha- and beta-carotene and lycopene determined by the HPLC. RESULTS: Serum alpha-carotene concentration did not change significantly in either of the experimental groups, whereas beta-carotene concentration decreased significantly in the WSEM and VOSEM groups (P<0.01), and the change differed significantly (P<0.05 and P<0.01, respectively) from that of the control group. Decrease in alpha+beta-carotene concentration was significantly greater (P<0.05) in both experimental groups than in the control group. However, the change

alpha-, beta- or alpha+beta-carotene/total cholesterol ratio did not differ significantly among the groups. No significant changes were found in serum lycopene or lycopene/total cholesterol ratio in both experimental

groups. CONCLUSIONS: Low-fat stanol ester margarines appeared to have little effect on serum concentrations of alpha-, beta- or alpha + beta-carotene, or lycopene. SPONSORSHIP: Grant to the University of Kuopio

by Raisio Benecol Ltd, Raisio, Finland.

L112 ANSWER 10 OF 40 MEDLINE

in

2000007614 Document Number: 20007614. Sitostanol administered in lecithin micelles potently reduces cholesterol absorption in humans. Ostlund R E Jr; Spilburg C A; Stenson W F. (Division of Endocrinology, Washington University, St Louis, MO, USA..

ROstlund@imgate.wustl.edu) . AMERICAN JOURNAL OF CLINICAL NUTRITION, Prepared by M. Hale 308-4258 Page 24

Nov) 70 (5) 826-31. Journal code: 3EY. ISSN: 0002-9165. Pub. country: United States. Language: English.

BACKGROUND: Phytosterol feeding in human clinical trials has had AB generally small and inconsistent effects on serum cholesterol concentrations, raising doubts about the importance of phytosterols in natural diets and supplements. OBJECTIVE: The hypothesis tested was that the low intestinal bioavailability of purified phytosterols can be increased by formulation with lecithin. DESIGN: The ability of sitostanol to reduce cholesterol absorption was measured directly by including hexadeuterated cholesterol tracer in a standard test breakfast and measuring plasma tracer concentration 4 and 5 d later by gas chromatography-negative ion mass spectrometry. The tracer amount after a test meal containing sitostanol was compared with that after an identical meal containing placebo. Each subject served as his or her own control and the order of testing was random. Sitostanol was formulated either as a powder or as a sonicated micellar solution with lecithin. A total of 38 single-meal tests were performed in 6 healthy subjects. RESULTS: Sitostanol powder (1 g) reduced cholesterol absorption by only 11.3 + - 7.4% (P = 0.2), confirming in vitro data showing poor solubility of sitostanol powder in artificial bile. In contrast, sitostanol in lecithin micelles reduced cholesterol absorption by 36.7 +/- 4.2% (P = 0.003) at a dose of 700 mg and by 34.4+/- 5.8% (P = 0.01) at a dose of 300 mg. CONCLUSIONS: Sitostanol reduced cholesterol absorption at doses lower than reported previously, but only if presented in lecithin micelles. Properly formulated sitostanol as well as naturally occurring complexes of phytosterol and phospholipid might be therapeutically useful for cholesterol lowering.

L112 ANSWER 11 OF 40 MEDLINE

1999208729 Document Number: 99208729. Serum sterols during stanol ester feeding in a mildly hypercholesterolemic population. Gylling H; Puska P; Vartiainen E; Miettinen T A. (Department of Medicine, University of Helsinki, P.O. Box 340, FIN-00029 HYKS, Helsinki, Finland.) JOURNAL OF LIPID RESEARCH, (1999 Apr) 40 (4) 593-600. Journal code: IX3. ISSN: 0022-2275. Pub. country: United States. Language: English.

AB We investigated the changes of cholesterol and non-cholesterol sterol metabolism during plant stanol ester margarine feeding in 153 hypercholesterolemic subjects. Rapeseed oil (canola oil) margarine without

(n = 51) and with (n = 102) stanol (2 or 3 g/day) ester was used for 1 year. Serum sterols were analyzed with gas-liquid chromatography. The latter showed a small increase in sitostanol peak during stanol ester margarine eating. Cholestanol, campesterol, and sitosterol proportions to cholesterol were significantly reduced by 5-39% (P < 0.05 or less for all) by stanol esters; the higher their baseline proportions the higher were their reductions. The precursor sterol proportions were significantly increased by 10- 46%, and their

baseline levels predicted low reduction of serum cholesterol. The decrease of the scheduled stanol dose from 3 to 2 g/day after 6-month feeding increased serum cholesterol by 5% (P < 0.001) and serum plant sterol proportions by 8-13% (P < 0.001), but had no consistent effect on precursor sterols. In twelve subjects, the 12-month level of LDL cholesterol exceeded that of baseline; Prepared by M. Hale 308-4258

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the non-cholesterol sterol proportions suggested that stimulated
synthesis
    with relatively weak absorption inhibition contributed to the
    non-responsiveness of these subjects. In conclusion, plant stanol ester
     feeding lowers serum cholesterol in about
     88% of subjects, decreases the non-cholesterol sterols that reflect
     cholesterol absorption, increases the sterols that reflect cholesterol
     synthesis, but also slightly increases serum plant stanols. Low synthesis
     and high absorption efficiency of cholesterol results in the greatest
    benefit from stanol ester consumption.
L112 ANSWER 12 OF 40 MEDLINE
1999268178 Document Number: 99268178.
                                         Cholesterol reduction by
     different plant stanol mixtures and with variable fat intake. Gylling H;
    Miettinen T A. (Department of Medicine, University of Helsinki, Finland.
)
    METABOLISM: CLINICAL AND EXPERIMENTAL, (1999 May) 48 (5) 575-80. Journal
     code: MUM. ISSN: 0026-0495. Pub. country: United States. Language:
     English.
AΒ
    Our aim was to investigate (1) whether different campestanol/
     sitostanol mixtures in margarine differ in reducing
     serum cholesterol, and (2) whether sitostanol
     ester in butter decreases serum cholesterol and alters
     cholesterol absorption and metabolism. Twenty-three postmenopausal women
     replaced 25 g dietary fat with (1) sitostanol ester-rich
     (campestanol to sitostanol ratio 1:11) and (2) campestanol
     ester-rich (campestanol to sitostanol ratio 1:2) rapeseed oil
    margarine, (3) butter, and (4) sitostanol ester-rich
     (campestanol to sitostanol ratio 1:13) butter. The respective
     scheduled stanol intake was 3.18, 3.16, and 2.43 g/d. The 6-week
margarine
     periods and, after an 8-week washout, 5-week butter periods were
     double-blind and in random order. Serum cholesterol
     precursor sterols (indicators of cholesterol synthesis) and plant sterols
     (indicators of cholesterol absorption) were quantified with gas-liquid
     chromatography (GLC). Low-density lipoprotein (LDL) cholesterol was
     reduced by 8% and 10% with the sitostanol and
     campestanol ester-rich margarines versus baseline (P < .05 for both) and
     high-density lipoprotein (HDL) cholesterol was increased by 6% and 5% (P
<
     .05), so the LDL/HDL cholesterol ratio was reduced by 15% (P <
     .05 for both). Sitostanol ester-rich butter decreased LDL
     cholesterol 12% and the LDL/HDL cholesterol ratio 11% (P < .05 for both)
     versus the butter period. The serum proportions of plant sterols and
     cholestanol were similarly reduced and those of cholesterol
     precursor sterols were similarly increased during all periods (P < .05
for
     all). Serum proportions of sitostanol and campestanol were
     slightly increased, indicating that their absorption related to their
     dietary intake. During all stanol interventions, serum vitamin D and
     retinol concentrations and alpha-tocopherol to cholesterol ratios were
     unchanged, whereas those of alpha- and beta-carotenes were significantly
     reduced. We conclude that varying the campestanol to
     sitostanol ratio from 1:13 to 1:2 in margarine and in butter
     similarly decreased cholesterol absorption, LDL cholesterol, and the
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LDL/HDL cholesterol ratio such that the serum lipids became less Prepared by M. Hale 308-4258

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atherogenic.

L112 ANSWER 13 OF 40 CAPLUS COPYRIGHT 2000 ACS 1999:244965 Document No. 130:295803 Sitostanol content in Setaria italica Beauv. seeds and milled grains. Abe, Bunichi; Itokawa, Emiko; Takatsuto, Suguru (Dep. Chem., Joetsu Univ. Educ., Joetsu, 943-8512, Japan). Nippon Nogei Kagaku Kaishi, 73(4), 419-421 (Japanese) 1999. CODEN: NNKKAA. ISSN: 0002-1407. Publisher: Nippon Nogei Kagakkai. From the nutritional point of view, it is known that sitostanol AB inhibits cholesterol absorption and lowers the level of serum cholesterol. The sitostanol content in Setaria italica Beauv. seeds and milled grain lipids was measured by GC anal. The result showed that each sample had a high content of sitostanol, ranging from 13 to 41 mg/100 g seeds or milled grains (27.6 mg, av., n = 9). No difference was found in **sitostanol** content between seeds and milled grains, suggesting that sitostanol is not localized in the aleurone layer, but distributed

L112 ANSWER 14 OF 40 MEDLINE

are good food materials.

1999173645 Document Number: 99173645. Effects of 2 low-fat stanol ester-containing margarines on **serum cholesterol** concentrations as part of a low-fat diet in hypercholesterolemic subjects.

uniformly in the seeds. It was also found that sitostanol

Hallikainen M A; Uusitupa M I. (Department of Clinical Nutrition, University of Kuopio, Finland. Maarit.Hallikainen@uku.fi) . AMERICAN JOURNAL OF CLINICAL NUTRITION, (1999 Mar) 69 (3) 403-10. Journal code: 3EY. ISSN: 0002-9165. Pub. country: United States. Language: English.

occurred mainly in free lipid. Nutritionally, S. italica milled grains

AB BACKGROUND: Full-fat sitostanol ester-containing margarine reduces serum total and LDL cholesterol, but the effect of plant stanol ester-containing margarine as part of a low-fat, low-cholesterol diet has not been studied. OBJECTIVE: We investigated the cholesterol-lowering effects of 2 novel, low-fat stanol ester-containing margarines as part of a low-fat diet recommended for hypercholesterolemic subjects. DESIGN: In a parallel, double-blind study, 55 hypercholesterolemic subjects were randomly assigned after a 4-wk high-fat

diet (baseline) to 3 low-fat margarine groups: wood stanol ester-containing margarine (WSEM), vegetable oil stanol ester-containing margarine (VOSEM), and control margarine (no stanol esters). The groups consumed the margarines for 8 wk as part of a diet resembling that of the National Cholesterol Education Program's Step II diet. The daily mean total stanol intake was 2.31 and 2.16 g in the WSEM and VOSEM groups, respectively. RESULTS: During the experimental period, the reduction in serum total cholesterol was 10.6% (P < 0.001) and 8.1% (P < 0.05) greater and in LDL cholesterol was 13.7% (P < 0.01) and 8.6% (P = 0.072) greater in the WSEM and VOSEM groups, respectively, than in the control group. Serum campesterol concentrations decreased 34.5%

and

41.3% (P < 0.001) in the WSEM and VOSEM groups, respectively. Serum HDL cholesterol, sitostanol, campestanol, beta-carotene, and fat-soluble vitamin concentrations did not change significantly from baseline. CONCLUSIONS: We conclude that the low-fat, plant stanol ester-containing margarines are effective cholesterol-lowering Prepared by M. Hale 308-4258

products in hypercholesterolemic subjects when used as part of a low-fat, low-cholesterol diet. They offer an additional, clinically significant reduction in serum cholesterol concentrations to that obtained with a low-fat diet alone.

L112 ANSWER 15 OF 40 MEDLINE

1999417057 Document Number: 99417057. Retinol, vitamin D, carotenes and alpha-tocopherol in serum of a moderately hypercholesterolemic population consuming sitostanol ester margarine. Gylling H; Puska P; Vartiainen E; Miettinen T A. (Department of Medicine, University of Helsinki, Finland.) ATHEROSCLEROSIS, (1999 Aug) 145 (2) 279-85. Journal code: 95X. ISSN: 0021-9150. Pub. country: Ireland. Language: English.

AB We have shown earlier that sitostanol ester margarine lowers serum cholesterol by inhibiting

cholesterol absorption so that, theoretically, there could be interference

with the absorption of fat-soluble vitamins. Accordingly, we investigated whether sitostanol ester margarine affects the serum levels of vitamin D, retinol, alpha-tocopherol and alpha- and beta-carotenes during 1-year treatment in 102 subjects and 49 controls with moderate hypercholesterolemia. The vitamins were assayed at baseline on home diet, on margarine alone, after 1 year's consumption of sitostanol ester margarine and after an additional 2 months on home diet. In the sitostanol group, serum plant sterols, indicators of cholesterol absorption efficiency, were reduced up to -38% in relation to controls from home diet (P < 0.01) indicating that cholesterol absorption was markedly reduced. Vitamin D and retinol concentrations and the ratio of alpha-tocopherol to cholesterol were unchanged by sitostanol ester. Serum beta-carotenes and alpha-carotene concentration but not proportion were reduced in the sitostanol group from baseline and in relation to controls (P < 0.01). Retinol and vitamin D were unassociated with serum cholesterol, plant sterols or other vitamins, whereas alpha-tocopherol and carotenes were significantly associated with serum plant sterols suggesting that the higher cholesterol absorption efficiency, the higher the alpha-tocopherol and carotene levels in serum. We conclude that sitostanol ester did not affect vitamin D and retinol concentrations and alpha-tocopherol/cholesterol proportion, but reduced serum beta-carotene levels. Alpha-tocopherol and carotenes, but not vitamin D and retinol, were related to serum cholesterol and cholesterol absorption.

L112 ANSWER 16 OF 40 CAPLUS COPYRIGHT 2000 ACS

1999:792367 Document No. 132:11900 Stanol esters in the **treatment** of hypercholesterolemia. Miettinen, T. A. (Department of Medicine, University of Helsinki, Helsinki, FIN-00029, Finland). Eur. Heart J. Suppl., 1(Suppl. S), S50-S57 (English) 1999. CODEN: EHJSFT. ISSN: 1520-765X. Publisher: W. B. Saunders Co. Ltd..

AB Aims: The extent and mechanisms of serum cholesterol lowering were studied by feeding mayonnaise and margarine without or with fortification with fat-sol. stanol esters to patients with mild hypercholesterolemia. Methods and Results: Double-blind, randomized, controlled studies were performed before and during feeding of mayonnaise or margarine with or without addn. of stanol esters to 242 mildly hypercholesterolemic participants. Serum lipids, ratios of non-cholesterol sterols to cholesterol, cholesterol absorption, and fecal Prepared by M. Hale 308-4258

elimination and synthesis of cholesterol were quantified. Replacement of home dietary fat with mayonnaise or margarine without stanol esters lowered serum total cholesterol levels by 5-9%. Total and low-d. lipoprotein (LDL) cholesterol levels were lowered up to 15% and 20%, resp., from home-diet values after feeding stanol ester margarine. Triglycerides and high-d. lipoprotein (HDL) cholesterol levels were unchanged. Cholesterol absorption efficiency and cholestanol and plant sterol ratios were decreased by up to 45% and resulted in proportionately enhanced elimination of cholesterol in stool as cholesterol but not as bile acids. Cholesterol synthesis and cholesterol precursor sterol

ratios

were increased compensatorily by more than 20%, a factor limiting the decrease of LDL cholesterol. Serum levels of vitamins A, E, D and .alpha.-carotene or ratios to cholesterol were not changed, but the .beta.-carotene ratio was decreased during 1-yr of stanol ester margarine feeding. Conclusion: Stanol ester margarine is a well-tolerated food ingredient that lowers serum total cholesterol levels by about 15% and LDL cholesterol levels by about 20% from home-diet values. Current hypocholesterolemic diets that include stanol ester margarine can normalize serum LDL cholesterol in more than half of participants; the rest may require combination treatment with statins.

L112 ANSWER 17 OF 40 CAPLUS COPYRIGHT 2000 ACS

1998:323105 Document No. 129:15518 Texturizing compositions for use in fat blends in food. Wester, Ingmar (Raisio Yhtyma Oy, Finland; Wester, Ingmar). PCT Int. Appl. WO 9819556 A1 19980514, 41 pp. DESIGNATED STATES: W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (English). CODEN: PIXXD2. APPLICATION: WO 1997-FI669 19971103. PRIORITY: US 1996-740845 19961104.

AB Fatty acid esters, such as the unsatd. fatty acid esters of sterols and/or

stanols, are used as a replacement for a substantial portion or all of the

undesirable satd. and trans-unsatd. fats used as structure giving hardstocks in edible foods such as margarines, mayonnaise, cooking oils, cheeses, butter and shortening. Because of the similarity in the crystallinity and phys. properties of the esters to those of the undesirable hardstock fats, the substitution or replacement contributes favorably to the flavor, texture and other sensory properties of the foods. Only the fatty acid portion of the phytosterol esters defined herein as texturizing agent is digested or absorbed with the sterol part being unabsorbable, thereby resulting in a redn. in total caloric uptake. Furthermore, the phytosterol fatty acid esters reduce the absorption of both dietary and biliary cholesterol from the digestive tract, thereby lowering the blood serum cholesterol level, esp. the LDL-cholesterol.

L112 ANSWER 18 OF 40 CAPLUS COPYRIGHT 2000 ACS

1998:124016 Document No. 128:191937 Stanol composition and the use thereof. Wester, Ingmar; Palmu, Tapio; Miettinen, Tatu; Gylling, Helena (Raision Tehtaat Oy AB, Finland; Wester, Ingmar; Palmu, Tapio; Miettinen, Tatu; Prepared by M. Hale 308-4258 Page 29

Gylling, Helena). PCT Int. Appl. WO 9806405 A1 19980219, 29 pp. DESIGNATED STATES: W: AT, AU, AZ, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK. EE, ES, FI, GB, GE, HU, IL, IS, JP, KE, KP, KR, KZ, LK, LT, LU, LV, MD, MK, MN, MX, NO, NZ, PL, PT, RO, RU, SE, SG, SI, SK, TJ, TM, TR, UA, US, UZ, VN, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (English). CODEN: PIXXD2. APPLICATION: WO 1996-FI465 19960902. PRIORITY: FI 1996-3126 19960809. AB A stanol compn. contg. in addn. to sitostanol as the main component, also a substantial amt. of at least 10 % campestanol effectively lowers serum cholesterol levels when incorporated in edible commodities. Upon esterification the compn. is esp. useful in edible fats and oils and in fat-contg. foods. vegetable oil distillate (brassicasterol 2.7, campesterol 26.7, stigmasterol 18.4, sitosterol 49.1, and sitostanol 2.9 %) was hydrogenated and esterified with rapeseed oil Me ester. Margarines

were produced using the above product.

- L112 ANSWER 19 OF 40 MEDLINE
- 1999058324 Document Number: 99058324. [Is the Finnish "healthy margarine" food or medicine? Addition of plant sterols can lower cholesterol levels]. Ar det finska "halsomargarinet" mat eller medicin? Tillsats av vaxtsteroler kan sanka hoga kolesterolvarden. Wikstrom A C. (Karolinska institutet, enheten for preventiv nutrition, Huddinge sjukhus.
 -) LAKARTIDNINGEN, (1998 Nov 11) 95 (46) 5146-8. Ref: 18. Journal code: LON. ISSN: 0023-7205. Pub. country: Sweden. Language: Swedish.
- AB Sine the autumn of 1995, Benecol, a proprietary brand of cholesterollowering margarine, has been available in ordinary grocery shops
 in Finland. The active ingredient is a sitostanol ester. Several
 studies in humans have shown use of the margarine to result in an
 approximately 10 per cent reduction in total serum
 cholesterol, and a 13-15 per cent reduction of
 LDL-cholesterol. However, further studies are required of its
 phyto-oestrogenic and endocrine effects, and its effects on growing
 children, particularly regarding subsequent fertility in boys. Although
 the margarine is classed as a 'functional food' in Finland, the question
 arises where the line is to be drawn between medicines and food-stuffs.
- L112 ANSWER 20 OF 40 CAPLUS COPYRIGHT 2000 ACS
 1999:52146 Document No. 130:261865 Serum cholesterol
 lowering effects of the phytosterol derivative (LPSS) in
 rats. Che, Jeong-Hwan; Chung, Dae-Won; Noh, Seung-Kwon; Lee, Yong-Soon;
 Park, Jae-Hak (Department of Veterinary Public Health, College of
 Veterinary Medicine, Seoul National University, Suwon, 441-744, S.
 Korea).
 - J. Toxicol. Public Health, 14(4), 535-539 (Korean) 1998. CODEN: JTPHFT. ISSN: 1226-8399. Publisher: Korean Society of Toxicology.
- AB The present study was designed to investigate the serum cholesterol lowering effect of the phytosterol deriv. (LPSS) on high cholesterol (HC) diet-induced hypercholesterolemia in male weaning Sprague-Dawley (SD) rats. Rats were fed with HC diet contg. 1% cholesterol and 0.5% cholic acid for 1 wk. After 1 wk, the LPSS

oil suspension (0.32 g/kg B.W.) was orally administered to the rats fed with either basal diet or HC diet groups for 7 days. In addn., the LPSS powder (0.14%) mixed with basal diet or HC diet was fed to the rats for 7 days. Serum total cholesterol and LDL-cholesterol contents were not altered by administration of the LPSS oil suspension with basal diet. However, they were significantly decreased by administration of the LPSS oil suspension with HC diet at day 14. Also, they were significantly decreased by the LPSS powder mixed with basal diet or HC diet at day 9, 11, 14. HDL-cholesterol contents were not altered by the LPSS oil suspension or LPSS powder. These results indicated that the phytosterol deriv.(LPSS) might decrease serum total cholesterol and LDL-cholesterol contents in rats.

- L112 ANSWER 21 OF 40 CAPLUS COPYRIGHT 2000 ACS
- 1998:636787 Document No. 130:24326 Effectivity and safety testing: Becel with **phytosterols** for effective cholesterol **lowering**.

 Louwes, A. C. M. (Neth.). Voeding, 59(9), 6-8 (Dutch) 1998. CODEN: VOEDAK. ISSN: 0042-7926. Publisher: Keesing Noordervliet.
- AB Becel margarine with various plant sterol additives was tested in volunteers at a dose of 3 g total sterols/day for its effectiveness in lowering serum cholesterol levels. Margarine contg. esterified soybean sterols lowered the total cholesterol and LDL cholesterol levels by 7-8 and 12-13%, resp.; this effect was dose dependent, and HDL cholesterol levels and other blood parameters remained unchanged. Margarine contg. sitostanol esters was somewhat less effective, and margarine contg. rice germ sterols or shea seed sterols had

little effect on cholesterol levels. All the margarines lowered the serum carotenoid levels. Toxicol. testing showed no mutagenic, pseudoestrogenic, hematol., or gastrointestinal effects of plant steroids.

- L112 ANSWER 22 OF 40 CAPLUS COPYRIGHT 2000 ACS
- 1997:617966 Document No. 127:257634 Z-2-[4-(4-Chloro-1,2-diphenyl-but-1-enyl)phenoxy]ethanol as serum cholesterol
 lowering agent, preparation thereof, and pharmaceutical
 compositions. Harkonen, Pirkko; Miettinen, Tatu; Mantyla, Eero; Kangas,
 Lauri; DeGregorio, Michael (Orion-Yhtyma Oy, Finland; Harkonen, Pirkko;
 Miettinen, Tatu; Mantyla, Eero; Kangas, Lauri; DeGregorio, Michael). PCT
 Int. Appl. WO 9732574 A1 19970912, 10 pp. DESIGNATED STATES: W: AM, AU,
 AZ, BA, BG, BR, BY, CA, CN, CZ, EE, FI, GE, HU, IL, IS, JP, KG, KP, KR,
 KZ, LK, LT, LV, MD, MK, MX, NO, NZ, PL, RO, RU, SG, SI, SK, TJ, TM, TR,
 UA, US, UZ, VN, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, CH, DE,
 DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE. (English). CODEN:
 PIXXD2. APPLICATION: WO 1997-FI140 19970304. PRIORITY: GB 1996-4577
 19960304.
- AB A method of lowering serum cholesterol levels comprises administering to a patient in need of such treatment an effective amt. of Z-2-[4-(4-chloro-1,2-diphenyl-but-1-enyl)phenoxy]ethanol (I). Pharmaceutical compns. useful in the method are
- also disclosed, as is use of I in the manuf. of a medicament for the prevention or treatment of atherosclerosis. Prepn. of I is described.
- L112 ANSWER 23 OF 40 MEDLINE Prepared by M. Hale 308-4258

1998077368 Document Number: 98077368. Reduction of serum cholesterol in postmenopausal women with previous myocardial infarction and cholesterol malabsorption induced by dietary sitostanol ester margarine: women and dietary sitostanol . Gylling H; Radhakrishnan R; Miettinen T A. (Department of Medicine, University of Helsinki, Finland.) CIRCULATION, (1997 Dec 16) 96 (12) 4226-31. Journal code: DAW. ISSN: 0009-7322. Pub. country: United States.

Language: English.

BACKGROUND: Reduction of serum cholesterol AB decreases mortality in primary and especially in secondary prevention. We investigated how effectively postmenopausal women with a previous myocardial infarction reduced their serum cholesterol with dietary means by using sitostanol ester rapeseed oil margarine, alone and in combination with statins, and to what

extent cholesterol metabolism was affected. METHODS AND RESULTS: The first

study group consisted of 22 randomly chosen women with angiographically documented coronary artery disease. Baseline studies on home diet were followed by double-blind, randomized, cross-over studies on margarine without and with sitostanol (3 q/d) ester for 7 weeks in random order. A second group of 10 women on simvastatin consumed sitostanol ester margarine for 12 weeks. Sitostanol ester margarine lowered serum total cholesterol by 13% (P<.05) and LDL cholesterol by 20% (P<.01). Sitostanol ester margarine reduced total cholesterol in all patients, LDL cholesterol <2.6 mmol/L (<100 mg/dL) in 32%, and <3.4 mmol/L (<133 mg/dL) in 73% versus none and 27% during the home diet (P<.01 for both). Combined with simvastatin, sitostanol still reduced total and LDL cholesterol by 11+/-3% and 16+/-5% (P<.01 for both). Sitostanol reduced absorption (-45%), increased fecal elimination (+45% as neutral sterols), and stimulated synthesis (+39%) of cholesterol. High cholestanol and plant sterol (high cholesterol absorption) and low baseline precursor sterol proportions (low cholesterol synthesis) predicted high decreases in serum cholesterol. CONCLUSIONS: Dietary use of sitostanol ester margarine normalizes LDL cholesterol in about one third of women with previous myocardial infarction, especially in those with high baseline absorption and low synthesis of cholesterol, and in combination with statins reduces the needed drug dose.

L112 ANSWER 24 OF 40 MEDLINE

96048304 Document Number: 96048304. Sterol absorption and sterol balance in phytosterolemia evaluated by deuterium-labeled sterols: effect of sitostanol treatment. Lutjohann D; Bjorkhem I; Beil U F; von Bergmann K. (Department of Clinical Pharmacology, University of Bonn, Germany..) JOURNAL OF LIPID RESEARCH, (1995 Aug) 36 (8) 1763-73. Journal

code: IX3. ISSN: 0022-2275. Pub. country: United States. Language: English.

AΒ Absorption of dietary cholesterol, campesterol, and sitosterol, cholesterol balance, and fecal excretion of plant sterols were determined in three unrelated patients with phytosterolemia and three healthy volunteers during constant intake of cholesterol and plant sterols Prepared by M. Hale 308-4258

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using accurate gas-liquid chromatography-mass spectrometry techniques. Each subject received a mixture of [26,26,26,27,27,27-2H6]cholesterol, [6,7,7-2H3] sitostanol, and [6,7,7-2H3] campesterol together with two non-absorbable markers, [5,6,22,23-2H4] sitostanol and chromic oxide. Feces were collected from days 5 to 7 and absorption of different sterols was calculated from the intestinal disappearance of the different sterols relative to [5,6,22,23-2H4] sitostanol and chromic oxide. The results obtained by the two markers were not different and the absorption of cholesterol averaged 53 +/- 4% for the patients (mean +/- SD) and 43 +/- 3% for the volunteers. Campesterol absorption averaged 24 +/- 4% in patients and 16 +/- 3% in healthy volunteers, whereas sitosterol absorption averaged 16 +/- 1% and 5 +/- 1%, respectively. Cholesterol synthesis expressed by body weight varied considerably in the two groups but appeared to be about 5 times lower in patients than in controls. Administration of a high dose of sitostanol (0.5 g t.i.d.) to two patients was followed by a reduction in cholesterol absorption by 24% and 44%, an increase in fecal output of cholesterol and steroids derived from cholesterol and plant steroids, and a marked reduction of serum cholesterol, campesterol, and sitosterol. Under the conditions used, inhibition of cholesterol absorption by sitostanol was not followed by a significant rise in cholesterol synthesis. The time of observation was, however, too short to allow final conclusion on this. The results show that the absolute difference in absorption rate of different sterols between the patients and healthy volunteers was about the same. As a consequence, increasing hydrophobicity

causes a relative decrease of absorption rates. Thus, patients with **phytosterolemia** seem to have a generally increased absorption of sterols rather than a loss of a specific discriminatory mechanism, and oral administration of **sitostanol** seems to be an interesting new approach for **treatment** of **phytosterolemia**.

L112 ANSWER 25 OF 40 MEDLINE

96036647 Document Number: 96036647. Reduction of serum cholesterol with sitostanol-ester margarine in a mildly hypercholesterolemic population [see comments]. Miettinen T A; Puska P; Gylling H; Vanhanen H; Vartiainen E. (Department of Medicine, University of Helsinki, Finland...) NEW ENGLAND JOURNAL OF MEDICINE, (1995 Nov 16) 333 (20) 1308-12. Journal code: NOW. ISSN: 0028-4793. Pub. country: United States. Language: English.

AB BACKGROUND. Dietary plant sterols, especially sitostanol, reduce serum cholesterol by inhibiting cholesterol absorption. Soluble sitostanol may be more effective than a less soluble preparation. We tested the tolerability and cholesterol-lowering effect of margarine containing sitostanol ester in a population with mild hypercholesterolemia. METHODS. We conducted a one-year, randomized, double-blind study in 153 randomly selected subjects with mild hypercholesterolemia. Fifty-one consumed margarine without sitostanol ester (the control group), and 102 consumed margarine containing sitostanol ester (1.8 or 2.6 g of sitostanol per day). RESULTS. The margarine containing sitostanol ester was well tolerated. The mean one-year reduction in serum cholesterol was 10.2 percent in the sitostanol group, as compared with an increase of 0.1 percent in the control group. The difference in the change in Prepared by M. Hale 308-4258

serum cholesterol concentration between the two groups was -24 mg per deciliter (95 percent confidence interval, -17 to -32; P < 0.001). The respective reductions in low-density lipoprotein (LDL) cholesterol were 14.1 percent in the sitostanol group and 1.1 percent in the control group. The difference in the change in LDL cholesterol concentration between the two groups was -21 mg per deciliter (95 percent confidence interval, -14 to -29; P < 0.001). Neither serum triglyceride nor high-density lipoprotein cholesterol concentrations were affected by sitostanol. Serum campesterol, a dietary plant sterol whose levels reflect cholesterol absorption, was decreased by 36 percent in the sitostanol group, and the reduction was directly correlated with the reduction in total cholesterol (r = 0.57, P < 0.001). CONCLUSIONS. Substituting sitostanol-ester margarine for part of the daily fat intake in subjects with mild hypercholesterolemia was effective in lowering serum total cholesterol and LDL cholesterol.

L112 ANSWER 26 OF 40 MEDLINE

DUPLICATE 3

- 96113514 Document Number: 96113514. The effect of cholesterol absorption inhibition on low density lipoprotein cholesterol level. Gylling H; Miettinen T A. (Second Department of Medicine, University of Helsinki, Finland.) ATHEROSCLEROSIS, (1995 Oct) 117 (2) 305-8. Journal code: 95X. ISSN: 0021-9150. Pub. country: Ireland. Language: English.
- The degree of serum cholesterol lowering by up to almost maximal inhibition of cholesterol absorption was tested during neomycin and neomycin + sitostanol treatment in six hypercholesterolemic men. Neomycin decreased cholesterol absorption efficiency by 49% and the combination by 79%, and serum cholesterol level by 27% and 36%, respectively. The correlation between the absorption percentage and low density lipoprotein (LDL) cholesterol was significant (r = 0.510), and the regression equation (y = 0.04x + 2.59) suggested that the mean LDL cholesterol content would be about 2.5 mmol/l at zero cholesterol absorption. In conclusion, in hypercholesterolemic subjects, the lowering of LDL cholesterol appears to be limited to a low normal range only by almost totally inhibiting cholesterol absorption.
- L112 ANSWER 27 OF 40 CAPLUS COPYRIGHT 2000 ACS
- 1995:639977 Document No. 123:65634 Serum cholesterollowering effects and triterpenoids of the herbs of Lactuca indica.
 Park, Hee Juhn; Lee, Myung Sun; Lee, Eun; Choi, Moo Young; Cha, Bae Chun;
 Jung, Won Tae; Young, Han Suk (Coll. Life Sci. Natural Resour., Sangji
 Univ., Wonju, 220-702, S. Korea). Saengyak Hakhoechi, 26(1), 40-6
 (Korean) 1995. CODEN: SYHJAM. ISSN: 0253-3073.
- AB A methanol ext. of the herbs of Lactuca indica L. effectively decreased the serum levels of total cholesterol and LDL-cholesterol when orally administered with diet. A chloroform-sol. fraction showed the similar effects with the methanol ext. Chromatog. sepn. afforded a mixt. of triterpene alcs. and their acyl derivs. A mixt. of triterpene alcs. were identified as .beta.-amyrin, .alpha.-amyrin, lupeol, pseudotaraxasterol, taraxasterol, and germanicol by spectroscopic methods. The acyl moieties in the corresponding acyl mixt. were characterized as acetates and palmitates. Three kinds of sterols such as .beta.-sitosterol, compesterol and stigmasterol were isolated as a mixt. state.

95163652 Document Number: 95163652. Cholesterol malabsorption caused by sitostanol ester feeding and neomycin in pravastatintreated hypercholesterolaemic patients. Vanhanen H. (Second
Department of Medicine, University of Helsinki, Finland..) EUROPEAN
JOURNAL OF CLINICAL PHARMACOLOGY, (1994) 47 (2) 169-76. Journal code:
EN4. ISSN: 0031-6970. Pub. country: GERMANY: Germany, Federal Republic

Language: English.

of.

- AB Serum cholesterol values were insufficiently reduced by pravastatin in two different patient populations. Therefore, we studied whether further cholesterol reduction could be achieved by inhibiting both cholesterol synthesis (by pravastatin) and absorption (by neomycin or sitostanol ester). Thus, we measured serum cholesterol, cholesterol precursors (reflecting cholesterol synthesis), cholestanol and plant sterols (reflecting cholesterol absorption and biliary secretion) for up to 6 weeks in pravastatin-treated patients with familial hypercholesterolaemia (FH, n = 13) and with and without ileal bypass during addition of neomycin (1.5 g per day) and in another patient population of non-FH (n = 14) subjects during addition of sitostanol ester (1.5 g per day). Addition of neomycin lowered serum total, LDL and HDL cholesterol by a further 20%, and increased the pravastatin-lowered precursor: cholesterol ratios by 20% (irrespective of ileal bypass). It also reduced by 20% the plant sterol:cholesterol ratio (irrespective of ileal bypass) which was markedly increased by pravastatin alone. Pravastatin and neomycin in combination lowered total, LDL and HDL cholesterol by 45%, 53% and 17%, respectively. This combined regimen reduced the serum lathosterol:cholesterol ratio to about half of the reduction caused by pravastatin, while the elevation of the plant sterols: cholesterol ratio was less with the combination than with pravastatin alone. Changes in serum cholesterol precursor: cholesterol and plant sterol: cholesterol ratios during the combined treatment were smaller in the subgroup with ileal bypass. Addition of sitostanol ester did not lower serum total or LDL cholesterol nor the precursor:cholesterol ratios significantly, while the reduction observed in the plant sterols: cholesterol ratios was similar to that achieved with neomycin addition. (ABSTRACT TRUNCATED AT 250 WORDS)
- L112 ANSWER 29 OF 40 CAPLUS COPYRIGHT 2000 ACS
 1994:671840 Document No. 121:271840 Effect of the acyl-CoA:cholesterol
 acyltransferase inhibitor DuP 128 on cholesterol absorption and
 serum cholesterol in humans. Hainer, James W.; Terry,
 J. Greg; Connell, Jill M.; Zyruk, Hanna; Jenkins, Rhonda M.; Shand, Donna
 L.; Gillies, Peter J.; Livak, Kenneth J.; Hunt, Thomas L.; et al. (DuPont
 Merck Pharmaceutical Company, Wilmington, DE, 19880-0026, USA). Clin.
 Pharmacol. Ther. (St. Louis), 56(1), 65-74 (English) 1994. CODEN:

ISSN: 0009-9236.

CLPTAT.

- AB Intestinal cholesterol esterification by the enzyme acyl-CoA:cholesterol acyltransferase (ACAT) is a presumed prerequisite for cholesterol absorption. We evaluated the effect of a potent, poorly absorbed ACAT inhibitor (DuP 128:
- N'-(2,4-difluorophenyl)-N-[5,(4,5-diphenyl-1H-imidazol-2-ylthio)pentyl]-N-heptylurea) on cholesterol absorption in a randomized Prepared by M. Hale 308-4258 Page 35

trial. Thirty subjects received DuP 128 for 7 wk, 10 each at 900 mg per day, 1800 mg per day, and 3600 mg per day; six subjects received placebo; and nine subjects received 1 gm neomycin twice a day. Cholesterol absorption detns. used a continuous dual isotope 14C-cholesterol and 3H-beta sitosterol method. DuP 128 (pooled doses) induced at 14.4% .+-. 11.4% redn. in cholesterol absorption (p < 0.05 vs. placebo): 17.6% .+-. 8.4% at 900 mg, 9.1% .+-. 11.4% at 1800 mg, and 17.1%

.+-. 12.9% at 3600 mg. Neomycin induced a 26.4% .+-. 10.7% redn . (p < 0.01). After 6 wk, neomycin reduced serum total and low-d. lipoprotein cholesterol by 22.4% .+-. 9.2% and 24.0% .+-. 11.6%, resp. (p < 0.01 vs. placebo). DuP 128 induced redns. of 3.9% .+-. 11% (difference not significant) and 4.95% .+-. 14.3% (p = 0.05). ACAT inhibitors limit cholesterol absorption in humans; however, the magnitude of the effect, as exemplified by DuP 128, is small.

L112 ANSWER 30 OF 40 MEDLINE

91298880 Document Number: 91298880. Hypocholesterolemic activity of betasitosterol in cholesterol fed sea quail. Day C E. (Audax, Inc.,
Leitchfield, KY 42754...) ARTERY, (1991) 18 (3) 125-32. Journal code:
8NN. ISSN: 0098-6127. Pub. country: United States. Language: English.
AB Male SEA quail were fed a 0.5% cholesterol supplemented diet, to which
was

added 0%, 1%, and 2% beta-sitosterol, for a period of seven days. Dietarily administered beta-sitosterol reduced total serum cholesterol levels by 62% and 72% at the 1% and 2% treatment doses, respectively. This hypocholesterolemic activity of sitosterol in cholesterol fed SEA quail was anticipated on the basis of the numerous earlier studies demonstrating similar activity in cholesterol fed chickens. Beta-sitosterol was tested in SEA quail to experimentally confirm its expected serum cholesterol lowering effects and to expand further the utility of the SEA quail model in cholesterol and atherosclerosis research.

L112 ANSWER 31 OF 40 BIOSIS COPYRIGHT 2000 BIOSIS

1990:429645 Document No.: BA90:90446. ON THE HYPOCHOLESTEREMIC

EFFECT OF ERYNGIUM-HETEROPHYLLUM. NAVARRETE A; NINO D; REYES B; SIXTOS C;

AGUIRRE E; ESTRADA E. LABORATORIO PRODUCTOS NATURALES, AREA DE QUIMICA,

DEP. PREPARATORIA AGRICOLA, UNIVERSIDAD AUTONOMA CHAPINGO, CHAPINGO,

ESTADO DE MEXICO, CP 56230, MEXICO.. FITOTERAPIA, (1990) 61 (2), 182-184.

CODEN: FTRPAE. ISSN: 0367-326X. Language: English.

AB Oral administration of an aqueous extract of E. heterophyllum caused reduction in the serum cholesterol in rats and significant hypotensive effect in humans. Mannitol, glucose and .beta.-sitosterol were identified as constituents of this plant.

L112 ANSWER 32 OF 40 MEDLINE

90032859 Document Number: 90032859. The mechanism of the hypocholesterolaemic effect of activated charcoal. Neuvonen P J; Kuusisto P; Manninen V; Vapaatalo H; Miettinen T A. (Department of Clinical Pharmacology, University of Helsinki, Finland..) EUROPEAN JOURNAL OF CLINICAL INVESTIGATION, (1989 Jun) 19 (3) 251-4. Journal code: EN3. ISSN:

0014-2972. Pub. country: ENGLAND: United Kingdom. Language: English.

AB The hypocholesterolaemic mechanism of activated charcoal was studied in Prepared by M. Hale 308-4258 Page 36

seven patients with primary hypercholesterolaemia. The reduction of serum cholesterol was correlated with the serum concentrations of cholesterol precursors and of two plant sterols. Activated charcoal, 8 g t.i.d. for 4 weeks, reduced serum concentration of total cholesterol by 27% (P less than 0.01). The effect was accompanied by a moderate elevation (P less than 0.05) in serum squalene and desmosterol concentrations and by a marked increase (up to 300-700%) in serum lathosterol and delta 8 lathosterol concentrations.

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levels of two plant sterols, campesterol and beta-sitosterol, were unchanged or only slightly decreased by the use of activated charcoal. The decrease of serum cholesterol

concentration had significant negative correlations with serum lathosterol

and delta 8 lathosterol, and significant positive correlations with serum cholestanol and beta-sitosterol. These observations suggest an increased cholesterol synthesis upon treatment with activated charcoal, probably caused by the interference with the enterohepatic circulation of bile acids.

- L112 ANSWER 33 OF 40 CAPLUS COPYRIGHT 2000 ACS
- 1984:208354 Document No. 100:208354 Hypocholesteremic effect of linoleic acid and phytosterol. I. Change in serum lipids in healthy men. Hasegawa, Kyoko; Shibuya, Kaori (Kagawa Nutr. Coll., Tokyo, Japan). Joshi Eiyo Daigaku Kiyo, 14, 165-72 (Japanese) 1983. CODEN: JEDKD7.
- AB Twenty men (av. age 36.6 yr) were divided into 4 groups and each group was
 - given 3, 2, 1, or 0 g cottonseed sterol (82% sitosterol [83-46-5]) day/person for 5 days, fed with trilinolein [537-40-6] (60 g) or butterfat (50 g) and sunflower oil (10 g). Linoleic acid [60-33-3] lowered the serum cholesterol [57-88-5] levels, probably by accelerating the catabolism of cholesterol to bile acid. Administration of plant sterol (mainly sitosterol) reduced absorption of cholesterol. Thus vegetable oils, esp. germ oils, have hypocholesteremic effects since they are high in linoleic acid and sitosterol.
- L112 ANSWER 34 OF 40 MEDLINE
- 82032551 Document Number: 82032551. Antihypercholesterolemic activity of beta-sitostanol in rabbits. Ikeda I; Kawasaki A; Samezima K; Sugano M. JOURNAL OF NUTRITIONAL SCIENCE AND VITAMINOLOGY, (1981) 27 (3) 243-51. Journal code: JFD. ISSN: 0301-4800. Pub. country: Japan. Language: English.
- AB The antihypercholesterolemic activity of beta-sitosterol and beta-sitostanol was compared in male rabbits given a cholesterol-supplemented diet. beta-Sitosterol and beta-sitostanol were fed to these rabbits at the 0.5% level with cholesterol (0.5% and 0.2% in experiments I and II, respectively). The serum cholesterol level tended to be lower in rabbits fed beta-sitostanol than in the animals fed beta-sitosterol even in experiment I. The beta-sitostanol exhibited a significantly greater hypocholesterolemic activity in experiment II, LDL-cholesterol being decreased markedly. The liver cholesterol decreased in both groups of rabbits to a similar extent.

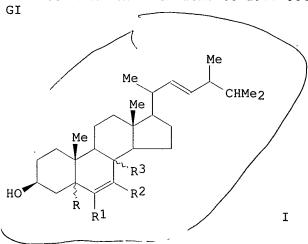
beta-

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Sitostanol prevented more effectively the formation of dietary cholesterol-induced atheroma in the abdominal aorta than beta-sitosterol. It is most likely, together with the data reported previously on rats, that the hypocholesterolemic activity of beta-sitostanol results from the significantly greater inhibitory effect on the intestinal absorption of cholesterol than that of beta-sitosterol.

L112 ANSWER 35 OF 40 CAPLUS COPYRIGHT 2000 ACS

1981:109362 Document No. 94:109362 Ergostadientriols and compositions containing them. Zilliken, Fritz W. (Z-L Ltd., USA). U.S. US 4234577 19801118, 7 pp. Cont.-in-part of U.S. 4,157,984. (English). CODEN: USXXAM. APPLICATION: US 1977-804594 19770608.



AB Pharmaceutical compns. contg. the title compds. I (R, R1, R2, and R3 = H or OH) are antioxidants and anticholesteremics. Thus, in an assay on chicks to det. the effect of I on serum cholesterol levels, I(R = R3 = OH, R1 = R2 = H) [76420-88-7] was 10 times as active as the known .beta.-sitosterol in reducing serum cholesterol. An antioxidant compn. also contained the isoflavone 2,3-dihydro-6,7-dihydroxy-3-(p-methoxyphenyl)-4H-1-benzopyran-4-one [76397-87-0].

L112 ANSWER 36 OF 40 MEDLINE DUPLICATE 4
78020099 Document Number: 78020099. Hypocholesterolemic action of a novel delta8-dihydroabietamide derivative, THD-341, in rats. Enomoto H;
Yoshikuni Y; Ozaki T; Zschocke R; Ohata K. ATHEROSCLEROSIS, (1977 Oct) 28
(2) 205-15. Journal code: 95X. ISSN: 0021-9150. Pub. country:
Netherlands. Language: English.

AB The hypocholesterolemic properties of THD-341, N-(2,6-dimethylphenyl)-delta8-dihydroabietamide, were studied in rats. THD-341 reduced serum cholesterol levels in cholesterol-cholate-fed rats at a concentration of less than 0.001% in the diet or an oral dose of less

than 3 mg/kg, once a day. When compared in terms of the 50% inhibitory dose for **serum cholesterol** elevation (ID 50%, % in diet), THD-341 (0.0008%) was comparable to D-thyroxine (0.0005%), more potent than estradiol (0.003%), and far more potent than clofibrate (0.2%), beta-**sitosterol** (0.8%), cholestyramine (2%), or Prepared by M. Hale 308-4258

nicotinic acid (3%). A daily intravenous injection of THD-341 was also effective (ID 50%: 7 mg/kg). THD-341 reduced serum and liver cholesterol in rats made hypercholesterolemic by 0.3% dietary thiouracil or 0.25% dietary cholate. Liver cholesteriol was more profoundly affected than the serum cholesterol. In normal rats, cholesterol was reduced in liver but not in serum. Its mechanism of action is unknown but the results suggest that THD-341 inhibits cholesterol absorption or re-absorption.

- L112 ANSWER 37 OF 40 CAPLUS COPYRIGHT 2000 ACS
- 1975:552376 Document No. 83:152376 Anticholesteremic lecithin-.beta.-sitosterol composition. Ritter, Kurt (Ritter, D., und Co., Ger.). Ger. Offen. DE 2400518 19750710, 6 pp. (German) CODEN: GWXXBX. APPLICATION: DE 1974-2400518 19740107.
- AB The prepn., useful for the lowering of serum cholesterol consists of lecithin and .beta.-sitosterol

 (I) [83-46-5] in the ratio of about 2:1. I addn. reduces the daily dosage, in the form of capsules or tablets, to about 1/3 of the normal required for lecithin alone, or to about 5 g to 10/day.
- L112 ANSWER 38 OF 40 CAPLUS COPYRIGHT 2000 ACS
- 1975:491179 Document No. 83:91179 Antilipemid agent based on a soybean oil fraction. Kaneda, Takashi; Tabata, Toshikazu Ger. Offen. DE 2334652 19750130, 40 pp. (German). CODEN: GWXXBX. APPLICATION: DE 1973-2334652 19730707.
- AB An unsaponifiable fraction from soybean oil, contg. about 45% phytosterols about 20% tocopherols, showed oral antilipemic activity in doses (1.2-1.8 g/day) that did not cause liver or kidney damage or other toxic effects. The fraction was prepd. from deodorized soybean oil distillate by esterifying fatty acids with MeOH and removing them by mol. distillation at 170-80.degree. and 20-50 mm Hg. The

was mixt. with silicic anhydride, a binder, and antioxidant, and a surfactant, powd., kneaded with an organic solvent to a granulate, and encapsulated. Administration of the fraction orally in capsule at 1.2 g/day to hyperlipidemic patients decreased the **serum cholesterol** level from an av. of 260 to 225.9 mg/dl and .beta.-lipoproteins from 592.3 to 527.1 mg/dl after 2 weeks.

- L112 ANSWER 39 OF 40 CAPLUS COPYRIGHT 2000 ACS
- 1973:79479 Document No. 78:79479 Serum cholesterol reducing agents. Rastogi, S. K. (G. R. Med. Coll., Gwalior, India). Curr. Med. Pract., 16(8), 333-5, 339 (English) 1972. CODEN: CMDPAW.
- AB A review with 7 refs. Serum cholesterol [57-88-5]lowering agents such as estrogens, thyroid hormones, nicotinic
 acid [59-67-6], atromid [8075-95-4], triparanol [78-41-1],
 sitosterol [83-46-5], and polyunsatd. vegetable oils are
 discussed.
- L112 ANSWER 40 OF 40 CAPLUS COPYRIGHT 2000 ACS
- .1974:43793 Document No. 80:43793 Dietary factors and drugs affecting fecal sterol excretion. Forman, Donald T.; Taylor, C. Bruce (Med. Sch., Northwest. Univ., Evanston, Ill., USA). Treat. Hyperlipidemic States, 376-86. Editor(s): Casdorph, Herman Richard. Thomas: Springfield, Ill. Prepared by M. Hale 308-4258

(English) 1971. CODEN: 27HXAF.

AB A review with 60 refs. on the lowering of serum cholesterol (I) [57-88-5] by .beta.-sitosterol [83-46-5], neomycin [1404-04-2], and hydrophilic colloids.

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